A

6/17-93

08504054 93214054

Mobilization of hematopoietic stem and progenitor cell subpopulations from the marrow to the blood of mice following cyclophosphamide and/or granulocyte colony-stimulating factor.

Neben S; Marcus K; Mauch P

Joint Center for Radiation Therapy, Harvard Medical School, Boston, MA 02115.

Blood (UNITED STATES) Apr 1 1993, 81 (7) p1960-7, ISSN 0006-4971 Journal Code: A8G

Languages: ENGLISH

Document type: JOURNAL ARTICLE

- end of record -

?

Display 1/3/2

08453069 93163069

Peripheral blood stem cell mobilization by cytokines.

Ahmed T; Wuest D; Ciavarella D

New York Medical College, Valhalla 10595.

J Clin Apheresis (UNITED STATES) 1992, 7 (3) p129-31, ISSN 0733-2459

Journal Code: HID Languages: ENGLISH

Document type: JOURNAL ARTICLE; REVIEW; REVIEW, TUTORIAL

- end of record -

?

Display 1/3/3

08392390 93102390

Peripheral blood stem cell mobilization after stem cell factor or G-CSF treatment: rapid enrichment for stem and progenitor cells using the CEPRATE immunoaffinity separation system.

Heimfeld S; Fogarty B; McGuire K; Williams S; Berenson RJ

CellPro, Incorporated, Bothell, Washington 98021.

Transplant Proc (UNITED STATES) Dec 1992, 24 (6) p2818, ISSN 0041-1345 Journal Code: WE9

Languages: ENGLISH

Document type: JOURNAL ARTICLE

- end of record -

?

Display 1/3/4

08095168 92233168

[The role of stem cell mobilization in the scope of autologous blood stem cell transplantation]

Die Rolle der Stammzell-Mobilisation im Rahmen der Autologen Blutstammzell-Transplantation (ABSZT).

Korbling M

M. D. Anderson Cancer Center, University of Texas, Houston.

Beitr Infusionther 1991, 28 p233-41, ISSN 1011-6974 Journal Code: BZI

Languages: GERMAN Summary Languages: ENGLISH

Document type: JOURNAL ARTICLE; REVIEW; REVIEW, TUTORIAL English Abstract

- end of record -

?

Display 1/3/5

07865980 92003980

Transplantation with blood stem cells. Zander AR; Lyding J; Bielack S Department of Hematology-Oncology, Universitatskrankenhaus Eppendorf, Hamburg, FRG. 1991, 17 (2) p301-9, ISSN 0340-4684 Blood Cells Journal Code: A8H Languages: ENGLISH Document type: JOURNAL ARTICLE; REVIEW; REVIEW, TUTORIAL - end of record -Display 1/3/6 07765556 91284556 Mobilization of haemopoietic stem cells by cyclophosphamide into the peripheral blood of patients with haematological malignancies. Shepherd KM; Charles P; Sage RE; Dale BM; Norman JE; Kotasek D; Gregg A; Futter J Department Haematology/Oncology, Queen Elizabeth Hospital, Woodville, Australia. 1991, 13 (1) p25-32, ISSN 0141-9854 Clin Lab Haematol Journal Code: DKF Languages: ENGLISH Document type: JOURNAL ARTICLE - end of record -? Display 1/3/7 07408512 90315512 The median daily increment of leukocytes during hematopoietic recovery the myeloid progenitor cell yield during leukapheresis in reflects children. Emminger W; Emminger-Schmidmeier W; Hocker P; Gerhartl C; Kundi M; Gadner St. Anna Children's Hospital, Vienna, Austria. Bone Marrow Transplant Jun 1990, 5 (6) p419-24, ISSN 0268-3369 Journal Code: BON Languages: ENGLISH Document type: JOURNAL ARTICLE - end of record -? Display 1/3/8 07292259 90199259 Therapeutic efficacy of autologous blood stem cell transplantation (ABSCT): the role of cytotoxic/cytokine stem cell mobilization. Korbling M; Haas R; Knauf W; Holle R; Hunstein W Institut f. Medizinische Biometrie, Heidelberg University, FRG. Bone Marrow Transplant Jan 1990, 5 Suppl 1 p39-40, ISSN 0268-3369 Journal Code: BON Languages: ENGLISH Document type: CLINICAL TRIAL; JOURNAL ARTICLE - end of record -Display 1/3/9

06054168 87028168

Complement split product C5a mediates the lipopolysaccharide-induced mobilization of CFU-s and haemopoietic progenitor cells, but not the mobilization induced by proteolytic enzymes.

Molendijk WJ; van Oudenaren A; van Dijk H; Daha MR; Benner R Cell Tissue Kinet Jul 1986, 19 (4) p407-17, ISSN 0008-8730 Journal Code: CQA

Languages: ENGLISH

```
nocument type: JOOKNHL HRIICLE
                                 - end of record -
      Display 1/3/10
04259195
           81087195
  Fyran copolymer: effect of molecular weight on stem cell mobilization in
mice.
  Zander AR; Spitzer G; Verma DS; Ginzbarg S; Dicke KA
  Biomedicine May 1980, 33 (3) p69-72, ISSN 0300-0893
                                                               Journal Code:
A5P
  Contract/Grant No.: CA-24770; CA-23077
  Languages: ENGLISH
  Document type: JOURNAL ARTICLE
                                 - end of record -
?
      Display 1/3/11
03321676
           77223676
  Mobilization of B and T lymphocytes and haemopoietic stem cells by
polymethacrylic acid and dextran sulphate.
  van der Ham AC; Benner R; Vos O
  Cell Tissue Kinet
                      Jul 1977, 10 (4) p387-97, ISSN 0008-8730
Journal Code: CQA
  Languages: ENGLISH
  Document type: JOURNAL ARTICLE
                                 - end of record -
?
      Display 1/3/12
02860675
           76041675
  Polymethacrylic acid: induction of lymphocytosis and tissue distribution.
  Ross WM; Martens AC; van Bekkum DW
  Cell Tissue Kinet
                      Sep 1975, 8 (5) p467-77, ISSN 0008-8730
Journal Code: CQA
  Languages: ENGLISH
  Document type: JOURNAL ARTICLE
Set
        Items
                Description
S1
           12
                STEM(W) CELL(W) (MOBILIZ? OR PERIPHERALIZ?)
S2
           55
                VERY (W) LATE (W) ANTIGEN (W) 4
S3
            0
                S2 AND STEM(W) CELL
S4
                VASCULAR (W) CELL (W) ADHESION (W) MOLECULE
          183
S5
                S4 AND STEM(W) CELL
            1
Ţ
      Display 5/3/1
08217769
           92355769
  Role of beta 1 and beta 2 integrins in the adhesion of human CD34hi stem
cells to bone marrow stroma.
  Teixido J; Hemler ME; Greenberger JS; Anklesaria P
  Department of Tumor Virology, Dana Farber Cancer Institute, Boston,
Massachusetts 02115.
                  Aug 1992, 90 (2) p358-67, ISSN 0021-9738
  J Clin Invest
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B10815

Contract/Grant No.: CA-39851; DE-08798; CA42368

Journal Code: HS7

Languages: ENGLISH

Document type: JOURNAL ARTICLE

9805466 BIOSIS Number: 44055466
PERIPHERAL BLOOD STEM CELL MOBILIZATION AFTER STEM CELL FACTOR OR G-CSF
TREATMENT RAPID ENRICHMENT FOR STEM AND PROGENITOR CELLS USING THE CEPRATE
IMMUNOAFFINITY SEPARATION SYSTEM

HEIMFELD S; FOGARTY B; MCGUIRE K; WILLIAMS S; BERENSON R J
CELLPRO INC., 22322 20TH AVE. SE, SUITE 100, BOTHELL, WASH. 98021.
FIRST INTERNATIONAL CONGRESS OF THE CELL TRANSPLANT SOCIETY, PITTSBURGH,
PENNSYLVANIA, USA, MAY 31-JUNE 3, 1992. TRANSPLANT PROC 24 (6). 1992.
2818. CODEN: TRPPA

Language: ENGLISH

Document Type: CONFERENCE PAPER

- end of record -

Display 1/3/5

(Item 5 from file: 5)

9769786 BIOSIS Number: 44019786

COMPARISON OF PERIPHERAL BLOOD STEM CELL MOBILIZATION BY THREE REGIMENS JANSSEN W E; ELFENBEIN G J; LEE C; SMILEE R; CARTER R; PACH M; LEPARC G F; SINALDI E; MASSARO P; ET AL

H. LEE MOFFITT CANCER CENT., UNIV. SOUTH FLA. COLL. MED., TAMPA, FLA. 45TH ANNUAL MEETING OF THE AMERICAN ASSOCIATION OF BLOOD BANKS, SAN FRANCISCO, CALIFORNIA, USA, NOVEMBER 7-12, 1992. TRANSFUSION (ARLINGT) 32 (8 SUPPL.). 1992. 43S. CODEN: TRANA

Language: ENGLISH

Document Type: CONFERENCE PAPER

- end of record -

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Display 1/3/6 (Item 6 from file: 5)

9347558 BIOSIS Number: 43092558

PERIPHERAL BLOOD STEM CELL MOBILIZATION AFTER STEM CELL FACTOR SCF ON G-CSF TREATMENT RAPID ENRICHMENT FOR STEM AND PROGENITOR CELLS USING THE CEPRATE IMMUNOAFFINITY SEPARATION SYSTEM

HEIMFELD S; ANDREWS R; ZSEBO K; FOGARTY B; MCGUIRE K; WILLIAMS S; BERENSON R

CELLPRO INC., BOTHELL, WASH.

XXI ANNUAL MEETING OF THE INTERNATIONAL SOCIETY FOR EXPERIMENTAL HEMATOLOGY, PROVIDENCE, RHODE ISLAND, USA, JULY 26-30, 1992. EXP HEMATOL (N Y) 20 (6). 1992. 748. CODEN: EXHMA

Language: ENGLISH

Document Type: CONFERENCE PAPER

- end of record -

?

Display 1/3/7 (Item 7 from file: 5)

8884735 BIOSIS Number: 42109735

PERIPHERAL BLOOD STEM CELL MOBILIZATION RAPID ENRICHMENT OF PROGENITOR CELLS USING A UNIQUE BIOTIN-AVIDIN IMMUNOAFFINITY SEPARATION SYSTEM HEIMFELD S; BENSINGER W; FOGARTY B; MCGUIRE K; WILLIAMS S; ZSEBO K; BERENSON R

CELLPRO INC., BOTHELL, WASH. 98021.

KEYSTONE SYMPOSIUM ON HEMATOPOIESIS, TAMARRON, COLORADO, USA, FEBRUARY 27-MARCH 5, 1992. J CELL BIOCHEM SUPPL 0 (16 PART C). 1992. 65. CODEN: JCBSD

Language: ENGLISH

Document Type: CONFERENCE PAPER

- end of record -

?

Display 1/3/8 (Item 8 from file: 5)

8871887 BIOSIS Number: 42096887

PERIPHERAL BLOOD STEM CELL MOBILIZATION RAPID ENRICHMENT OF PROGENITOR

CELLS USING A UNIQUE BIUTIN AVIDIN IMMUNUAFFINITY SEPARATION SYSTEM BERENSON R; ANDREWS R; BENSINGER W; FOGARTY B; MCGUIRE K; WILLIAMS S; HEIMFELD S CELLPRO INC., BOTHELL, WASHINGTON 98021, USA. KEYSTONE SYMPOSIUM ON BONE MARROW TRANSPLANTATION, KEYSTONE, COLORADO, USA, JANUARY 19-26, 1992. J CELL BIOCHEM SUPPL Ø (16 PART A). 1992. 189. CODEN: JCBSD Language: ENGLISH Document Type: CONFERENCE PAPER - end of record -? Display 1/3/12 (Item 12 from file: 5) BIOSIS Number: 38109914 7329393 THERAPEUTIC EFFICACY OF AUTOLOGOUS BLOOD STEM CELL TRANSPLANTATION ABSCT THE ROLE OF CYTOTOXIC-CYTOKINE STEM CELL MOBILIZATION KOERBLING M; HAAS R; KNAUF W; HOLLE R; HUNSTEIN W MED. KLINIK POLIKLINIK V, INST. F. MED. BIOMETRIC, HEIDELBERG UNIV., FRG. INTERNATIONAL SYMPOSIUM ON PERIPHERAL BLOOD STEM CELL AUTOGRAFTS, MULHOUSE, FRANCE, OCTOBER 15-17, 1989. BONE MARROW TRANSPLANT 5 (SUPPL. 1). 1990. 39-40. CODEN: BMTRE Language: ENGLISH Document Type: CONFERENCE PAPER - end of record -Display 1/3/13 (Item 13 from file: 5) BIOSIS Number: 38109906 7329385 APPROACHES TO BLOOD STEM CELL MOBILIZATION INITIAL AUSTRALIAN CLINICAL JUTTNER C A; TO L B; HAYLOCK D N; DYSON P G; BRADSTOCK K F; DALE B M; ENNO A; SAGE R E; SZER J; TOOGOOD I R G LEUKAEMIA RES. UNIT, IMVS, FROME RD., ADELAIDE, SOUTH AUSTRALIA, AUST. 5000. INTERNATIONAL SYMPOSIUM ON PERIPHERAL BLOOD STEM CELL AUTOGRAFTS, MULHOUSE, FRANCE, OCTOBER 15-17, 1989. BONE MARROW TRANSPLANT 5 (SUPPL. 1). CODEN: BMTRE 1990. 22-24. Language: ENGLISH Document Type: CONFERENCE PAPER - end of record -(Item 16 from file: 5) Display 1/3/16 2965749 BIOSIS Number: 69003156 THE MECHANISM OF HEMOPOIETIC STEM CELL MOBILIZATION A ROLE OF THE COMPLEMENT SYSTEM WILSCHUT I J C; ERKENS-VERSLUIS M E; PLOEMACHER R E; BENNER R; VOS O DEP. CELL BIOL. GENET., ERASMUS UNIV., P.O. BOX 1738, ROTTERDAM, NETH. CELL TISSUE KINET 12 (3). 1979. 299-312. CODEN: CTKIA Full Journal Title: Cell and Tissue Kinetics Language: ENGLISH

- end of display -

Items Description Set STEM(W) CELL(W) (MOBILIZ? OR PERIPHERALIZ?) S1 19 S2 38 VERY (W) LATE (W) ANTIGEN (W) 4 S3 0 S2 AND STEM(W) CELL 252 VASCULAR (W) CELL (W) ADHESION (W) MOLECULE S4 **S**5 0 S4 AND STEM(W) CELL **S6** 0 S1 AND S2 C1 OND CA

?ds

Display 9/3/15 (Item 15 from file: 5)

7754915 BIOSIS Number: 90122915

VASCULAR CELL ADHESION MOLECULE-1 MEDIATES LYMPHOCYTE ADHERENCE TO

CYTOKINE-ACTIVATED CULTURED HUMAN ENDOTHELIAL CELLS

CARLOS T M; SCHWARTZ B R; KOVACH N L; YEE E; ROSSO M; OSBORN L; CHI-ROSSO G; NEWMAN B; LOBB R; HARLAN J M

DIV. HEMATOL., ZA-34, HARBORVIEW MED. CENT., 325 9TH AVE., SEATTLE, WASH. 98104.

BLOOD 76 (5). 1990. 965-970. CODEN: BLOOA

Full Journal Title: Blood

Language: ENGLISH



- ≕> d 1-5
- 1. 5,188,959, Feb. 23, 1993, Extracellular matrix protein adherent T cells; Allan B. Haberman, 435/240.243, 4, 240.2 CIMAGE AVAILABLEJ
- 2. 5,187,193, Feb. 16, 1993, Method for stimulating transplanted bone marrow cells; Richard F. Borch, et al., 514/476 [IMAGE AVAILABLE]
- 3. 5,169,765, Dec. 8, 1992, Method for stimulating production of bone marrow cell growth factors using dithiocarbamates; Richard F. Borch, et al., 435/70.4, 69.5, 69.52 CIMAGE AVAILABLEI
- 4. 4,965,195, Oct. 23, 1990, Interleukin-7; Anthony E. Namen, et al., 435/69.52, 91, 172.1, 172.3, 320.1; 530/350, 351; 536/24.3, 24.31 CIMAGE AVAILABLE]
- 5) 4,808,402, Feb. 28, 1989, Method and compositions for modulating neovascularization; Samuel J. Leibovich, et al., 424/78.06, 423, 618; 514/2 [IMAGE AVAILABLE]
- (FILE 'USPAT' ENTERED AT 11:27:00 ON 17 JUN 93)
 L1 58124 S BLOOD
 L2 0 S STEM(W)CELL#(W)(MOBILIZ? OR PERIPHERALIZ?)
 L3 0 S STEM(W)CELL#(W)(MOBILIZ?)
 L4 145 S STEM(W)CELL
 L5 154 S CYTOKINE
 L6 5 S L1 AND L4 AND L5
- => d 18 1-3

∷ >

- 1. 5,188,959, Feb. 23, 1993, Extracellular matrix protein adherent T cells; Allan B. Haberman, 435/240.243, 4, 240.2 [IMAGE AVAILABLE]
- 2. 4,965,195, Oct. 23, 1990, Interleukin-7; Anthony E. Namen, et al., 435/69.52, 91, 172.1, 172.3, 320.1; 530/350, 351; 536/24.3, 24.31 CIMAGE AVAILABLEJ
- 3. 4,808,402, Feb. 28, 1989, Method and compositions for modulating neovascularization; Samuel J. Leibovich, et al., 424/78.06, 423, 618; 514/2 [IMAGE AVAILABLE]

```
Seq. 1
0| | 0 IntelliGenetics
>0 <
FastDB - Fast Pairwise Comparison of Sequences
Release 5.4
Results file us-07-977-702-1-ngs.res made by shears on Tue 15 Jun 93 11:23:42-PDT.
Query sequence being compared:US-07-977-702-1 (1-360)
Number of sequences searched:
                                             25646
Number of scores above cutoff:
                                              4455
      Results of the initial comparison of US-07-977-702-1 (1-360) with:
   Data bank : N-GeneSeq 10, all entries
 10000--
U 5000- *
M
В
R
0
F 1000-
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SCORE 0		•	•		•	•	172	506	240	275	309	
GTDFU	Λ	2	7	Δ	5	 Q 0						

PARAMETERS

Similarity matrix U	nitary	K-tuple	4
Mismatch penalty	1	Joining penalty	30
Gap penalty	1.00	Window size	32
Gap size penalty	0.33		
Cutoff score	0		
Randomization group	0		
Initial scores to save	40	Alignments to save	15
Optimized scores to save	0	Display context	50

SEARCH STATISTICS

Scores:	Mean	Median	Standard Deviation
	17	14	13.65
Times:	CPU 00:03:12.02		Total Elapsed

Number of residues: 14371384
Number of sequences searched: 25646
Number of scores above cutoff: 4455

Cut-off raised to 9. Cut-off raised to 15. Cut-off raised to 22. Cut-off raised to 28. Cut-off raised to 32.

The scores below are sorted by initial score. Significance is calculated based on initial score.

A 100% identical sequence to the query sequence was not found.

The list of best scores is:

Sequence Name	Description	Init Length Scor	. Opt. e Score	Sig.	Frame
	*** 21 standard deviations	above mean #	***		
1. 011098	Encodes gamma heavy chain of			21.40	0
	**** 18 standard deviations	above mean #	***		
2. @10381	Chimeric MAb 9.2.27 heavy cha	363 27	0 279	18.54	0
	**** 17 standard deviations	above mean #	***		
3. @08607	Co-1 Heavy Chain V Region (mo	471 25	9 265	17.73	0
4. 022736	RSV19 VH.	348 25	6 299	17.51	0
5. @28739	cDNA of VH425 antibody cloned	501 25	6 269	17.51	0
	*** 16 standard deviations	above mean #	***		
6. 023863	ScFvB18 construct mutant #6.	770 23	6 280	16.05	0
7. 023862	ScFvB18 construct mutant #5.	770 23	6 280	16.05	0
8. 023861	ScFvB18 construct mutant #4.	770 23	6 280	16.05	0
9. 023860	ScFvB18 construct mutant #3.	770 23	6 280	16.05	0
10. 023859	ScFvB18 construct mutant #2.	770 23	6 280	16.05	0

```
actible construct.
                                                      770
                                                            236
                                                                  280
                                                                       16.05
                     **** 15 standard deviations above mean ****
  12. 023858
                    ScFvB18 construct mutant #1.
                                                            234
                                                                  279
                                                      770
                                                                       15.90
  13. 015164
                   VH186 region of anti-nitrophe
                                                      458
                                                            233
                                                                  277
                                                                       15.83
  14. 906227
                   VH domain of antibody C again
                                                      345
                                                            229
                                                                  245
                                                                       15.53
  15. N91482
                   Genomic to cDNA junction (V47
                                                      349
                                                            228
                                                                  269
                                                                       15.46
  16. 012013
                   Sequence encoding mouse MAb 2
                                                      477
                                                            226
                                                                  259
                                                                       15.31
                   Sequence encoding heavy chain
  17. 012057
                                                      477
                                                            226
                                                                  259
                                                                       15.31
  18. 927141
                   ICAM-1 inhibiting peptide 2.
                                                      387
                                                            225
                                                                  263
                                                                       15.24
  19. 006957
                   Genomic sequence encoding hea
                                                     2675
                                                            225
                                                                  260
                                                                       15.24
  20. N91820
                   DNA sequence of the V and J r
                                                      443
                                                            224
                                                                  262 15.17
  21. 028263
                   Fv(FRP51)-ETA fusion gene.
                                                     2012
                                                            223
                                                                  270
                                                                       15.09
  22. 028260
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                                                      748
                                                            222
                                                                  269
                                                                       15.02
  23. 012637
                   Monoclonal antibody DK3T heav
                                                     1570
                                                            222
                                                                  268
                                                                       15.02
                    **** 14 standard deviations above mean ****
  24. 028258
                   FWP51 heavy chain variable do
                                                      342
                                                            218
                                                                  259
                                                                       14.73
  25. N91146
                   2H7 Vh sequence.
                                                      458
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                                                                  266
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  26. N70968
                   Sequence of the anti-hepatiti
                                                      420
                                                            217
                                                                  264
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                                                                  264 14.65
  28. 004261
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  29. N70971
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                                                      459
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  30. 023342
                   Murine heavy chain variable r
                                                      354
                                                            216
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                                                                       14.58
  31. N91645
                   Heavy chain of monoclonal ant
                                                      540
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                                                                  270 14.58
  32. 004040
                   Anti-Leu 3a heavy chain varia
                                                            216
                                                                  262 14.58
                                                      765
  33. 004262
                   Encodes Colon Cancer monoclon
                                                      416
                                                            215
                                                                  239 14.51
  34. 008609
                   ME4 Heavy Chain V Region (mou
                                                      492
                                                            215
                                                                  261
                                                                       14.51
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                                                            214
                   Heavy chain variable region o
                                                      361
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                                                                       14.43
  36. 004258
                   Sequence homologous to TAG72
                                                      982
                                                            214
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                                                                      14.43
  37. 028522
                   Hypercalcaemia agent cDNA por
                                                      356
                                                            213
                                                                  251
                                                                       14.36
  38. 024790
                   Anti-Tac antibody heavy chain
                                                      433
                                                            213
                                                                  247 14.36
  39. 005600
                   Anti-Tac heavy chain variable
                                                      433
                                                            213
                                                                  247 14.36
  40. 005554
                   Sequence encoding variable re
                                                      446
                                                            213
                                                                  233 14.36
1. US-07-977-702-1 (1-360)
   011098
                Encodes gamma heavy chain of T84.66 monoclonal ant
      Q11098 standard; DNA; 861 BP.
 ID
 AC
      011098;
 DT
      08-MAY-1991 (first entry)
      Encodes gamma heavy chain of T84.66 monoclonal antibody.
 K₩
      MAb T84.66; gamma heavy chain; carcinoembryonic antigen; CEA;
 KW
      human adenocarcinoma; mouse-human chimaeric antibody; ss.
 OS
      Mus musculus.
 FH
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                      Location/Qualifiers
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     exon
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FT
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FT
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     /note= "putative"
FT
     promoter
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FT
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     /note= "putative"
FT
     promoter
                      169..176
FT
     /*tag= g
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     /note= "putative"
PN
     WD9101990-A.
PD
     21-FEB-1991.
```

19-JUL-1990; U04049.

PF

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PA
     (CITY ) CITY OF HOPE.
 PΙ
     Shively JE, Riggs AD, Neumaier M;
 DR
     WPI; 91-073486/10.
 DR
     P-PSDB; R11384.
 PT
     Novel anti-CEA antibody - comparable to ATCC Accession No. BH
 PT
     8747, produced by recombinant DNA, used in diagnosis of tumours
PS
     Claim 4; Page 18; 24pp; English.
CC
     The heavy chain variable region of murine MAb 84.66 was cloned and
CC
     sequenced as follows: Hybridoma DNA was extracted, completely
CC
     restricted with EcoRI and run on a gel. Fragments were extracted and
CC
     ligated in the EcoRI site of Lambda-ZAP.Phage were packaged and plated.
CC
     Plaque screening was with a 991bp XbaI fragment from the mouse
CC
     enhancer region, a 1.5kb cDNA fragment from the heavy chain
CC
     constant region gene of hybridoma CEA.66-E3 and a 5.4kb EcoRI
CC
     fragment containing an aberrantly rearranged heavy chain from
CC
     Sp2/0. Positive clones were further characterised by hybridisation
CC
     to J-region oligonucleotides (see @10842-@10846) and sequenced. The
CC
     murine gamma variable region gene was used to produce mouse V-human C
CC
     antibodies with high affinity for CEA. Chimaeric murine-human anti-
CC
     CEA Abs are used to diagnose human colon adenocarcinomas.
CC
     See also 010834-010841 and 010847-8.
SQ
     Sequence
              861 BP;
                        213 A;
                                          203 G;
                                                   237 T;
Initial Score
                   309 Optimized Score =
                                           316 Significance = 21.40
Residue Identity =
                   88% Matches
                                           322 Mismatches
                                                                33
Gaps
                    10 Conservative Substitutions
                                                                0
                                                       10
                                                                20
                                                GTCAAACTGCAGCAGTCTGGGG
                                                   1 1111111111111111
   GTGACAGTGGCAATCACTTTGCCTTTCTTTCTACAGGGGTCAATTCAGAGGTTCAGCTGCAGCAGTCTGGGG
 360
          370
                   380
                            390
                                              410
                                                       420
                                                               430
         30
                          50
                                   60
                                            70
                                                     80
                                                              90
   CAGAGCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCT
   CAGAGCTTGTGGAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCT
         440
                 450
                          460
                                   470
                                            480
                                                     490
                                                              500
      100
              110
                        120
                                 130
                                         140
                                                  150
                                                           160
   ATATGCACTGGGTGAAGCAGAGCCTGAACAGGGCCTGGAGTGGATTGGAAGGATTGATCCTGCGAGTGGCG
   ATATGCACTGGGTGAAGCAGAGGCCTGAACAGGGCCTGGAATGGATTGGAAGGATTGATCCTGCGAATGGTA
       510
               520
                        530
                                 540
                                          550
                                                   560
                                                            570
    170
             180
                      190
                               200
                                        210
                                                220
                                                         230
   ATACTAAATATGACCCGAAGTTCCAGGTCAAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGC
   580
              590
                       600
                               610
                                        620
                                                 630
                                                          640
  240
           250
                    260
                             270
                                      280
                                               290
                                                       300
   TGCAGCTCAGCAGCCTGACATCTGAGGACACTGCCGTCTACTACTGTGCAGACGGAATGTGGGTATCAACG-
   Ш
                                                      1 111 11 1 111
   TGCAGCTCACCAGCCTGACATCTGAGGACACTGCCGTCTATTATTGTGC-TCCG---TTTGGTTA-CTACGT
   650
            660
                     670
                              680
                                       690
                                                700
                                                             710
     310
              320
                       330
                               340
                                        350
   ----GGATATGCTCTGGACTTCTGGGGCCAAGGGACCACGGTCACCGTCTCCTCA
         GTCTGACTATGCTATGGCCTACTGGGGTCAAGGAACCTCAGTCACCGTCTCCTCAGGTAAGAATGGCCTCTC
      720
              730
                       740
                                                  770
                                750
                                         760
                                                           780
```

CAGGTCTTTATTTTTAACCTTTGTATGGACTTT

810

800

790

ΓN

TO-AAF-1101: A2-7071AE

```
2. US-07-977-702-1 (1-360)
  010381
              Chimeric MAb 9.2.27 heavy chain variable region DN
 ID
     910381 standard; DNA; 363 BP.
 AC
     010381;
     15-APR-1991 (first entry)
 DT
DE
     Chimeric MAb 9.2.27 heavy chain variable region DNA sequence.
KW
     Chimeric antibodies; human glycoprotein antigen; melanoma; cancer;
ΚĦ
05
     Mus musculus.
FH
     Keu
                   Location/Qualifiers
FT
     CDS
                   1..363
FT
     /*taq= a
FT
     /product= H-chain V-region of MAb 9.2.27
PN
     EP-411893-A.
PD
     06-FEB-1991.
PF
     31-JUL-1990; 308402.
PR
     31-JUL-1989; US-387665.
PA
     (ELIL ) ELI LILLY & CO.
PI
     Beavers LS, Bumol TF, Gadski RA;
DR
     WPI; 91-038771/06.
DR
     P-PSDB; R10541.
PT
     Monoclonal antibody contg. recombinant DNA - binds to human
PT
     chondroitin sulphate proteoglycan on melanoma cells for melanoma
PT
     treatment and diagnosis
PS
     Claim 11; page 18; 33pp; English.
CC
     This sequence, contained in vector pG4G21, is ligated to a DNA
CC
     sequence encoding a human heavy (H) chain constant (C) region, in
CC
     the construction of pN9.2.27G1. This vector is used to transform
CC
     host cells, in conjunction with vector pG9.2.27K contg. murine
CC
     light (L) chain variable (V) region and human L- chain C- region.
CC
     The resulting host cell expresses the chimeric antibody 9.2.27
CC
     which is directed against proteoglycans of human melanoma cells.
CC
     This chimeric monoclonal antibody (MAb) is useful in the diagnosis
CC
     and treatment of melanoma. The use of human C-regions avoids the
CC
     problems associated with murine monoclonals e.g. rapid clearance
CC
     from the bloodstream due to anti-self recognition.
CC
     See also @10379-81 and @10382-84.
59
     Sequence
               363 BP;
                         88 A;
                                 87 C;
                                         105 G;
                                                  83 T;
Initial Score
                   270 Optimized Score =
                                            279
                                                Significance = 18.54
Residue Identity =
                   78% Matches
                                       =
                                            285
                                                Mismatches
Gaps
                     6 Conservative Substitutions
                                                            =
                                                                  0
      X
                      20
             10
                               30
                                         40
                                                 50
                                                           60
      GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCT
      CAGGTCCAGCTGCAGCAGTCTGGACCTGAGCTGGTGAAGCCTGGGGCCTCAGTGAAGATTTCCTGCAAAGCT
                   20
      X
          10
                             30
                                      40
                                               50
                                                        60
 70
                     90
                             100
                                      110
                                                120
   TCTGGCTTCAACATTAAAGA-CACCTATATGCACTGGGTGAAGCAGAGCCTGAACAGGGCCTGGAGTGGAT
   TCTGGCTAC-GCATTCAGTAGGTCTTGGATGAACTGGGTGAAGCAGAGGCCTGGACAGGGTCTTGAGTGGAT
         80
                   90
                           100
                                    110
                                             120
                                                      130
                                                               140
         150
                             170
                                               190
                   160
                                      180
                                                        200
                                                                 210
   TGGAAGGATTGATCCTGCGAG-TGGCGATACTAAATATGACCCGAAGTTCCAGGTCAAGGCCACTATTACAG
   150
                160
                          170
                                   180
                                            190
                                                     200
                                                              210
         220
                  230
                           240
                                    250
                                             260
                                                      270
```

CGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGACACTGCCGTCTACTACT

```
......
                              CAGACAAATCCTCCAGCACAGCCTACATGCAGGTCAGCAGCCTGACCTCTGTGGACTCTGCGGTCTATTTCT
      220
                230
                          240
                                    250
                                              260
                                                        270
                                                                  280
        290
                  300
                            310
                                      320
                                                330
                                                          340
                                                                    350
   GTGC-AGACGGAATGTGGGTATCAACGGGATATGCTCTGGACTTCTGGGGCCAAGGGACCACGGTCACCGTC
                    1111 1
                                 *** ** ***** ***** ***** *** * ******
   GTGCAAGAGGGAAT-ACGGTAGTAGTTCCCTATACTATGGACTACTGGGGTCAAGGAACCTCAGTCACCGTC
    290
              300
                         310
                                   320
                                             330
      360
   TCCTCA
    111111
   TCCTCA
   360 X
3. US-07-977-702-1 (1-360)
   008607
               Co-1 Heavy Chain V Region (mouse).
 ID
     008607 standard; DNA; 471 BP.
AC
     008607;
     04-MAR-1993 (first entry)
DT
DE
     Co-1 Heavy Chain V Region (mouse).
KW
     Monoclonal antibody; chimera; light; heavy; chain; constant;
KW
     variable; antigen; diagnosis; cancer; tumour; ss.
OS
     Mus ausculus.
FH
     Keu
                     Location/Qualifiers
FT
     CDS
                     52..471
FT
     /*tag= a
PN
     ₩09002569-A.
PD
     22-MAR-1990.
PF
     06-SEP-1989; U03852.
PR
     06-SEP-1988; US-240624.
PR
     08-SEP-1988; US-241744.
PR
     13-SEP-1988; US-243739.
PR
     04-0CT-1988; US-253002.
PR
     19-JUN-1989; US-367641.
PR
     21-JUL-1989; US-382768.
PA
     (ITGE-) INT GENETIC ENG INC.
     Better MD, Horwitz AH, Robinson RR, Lei S, Chang CP;
PI
DR
     WPI; 90-115825/15.
DR
     P-PSDB; R09425.
PT
     Chimeric mouse-human antibodies - prepd. using genes coding for
PT
     constant human region murine variable region, esp. to 3 tumour
PT
PS
     Claim 13; Page 123 + Fig 22; 173pp; English.
CC
     Shown is the nucleotide sequence from the end of the oligo-dC tail
CC
     to the Jh4-Ch1 junction. The sequence is used in the prodn. of a
CC
     chimeric antibody mol. comprising two light chains and two heavy chains,
CC
     each having a constant region (human) and a variable region (murine)
CC
     having specificity to an antigen bound by murine monoclonal antibody
CC
     (MAb) Co-1. The chimeric antibodies can be used for any purpose for
CC
     which the original murine MAbs can be used, with the advantage that
     they are more compatible with the human body. They are esp. used for
CC
CC
     the diagnosis and treatment of cancer.
SQ
     Sequence 471 BP;
                           115 A;
                                     120 C;
                                               115 G;
                                                         121 T;
Initial Score
                     259 Optimized Score =
                                                265 Significance = 17.73
Residue Identity = .
                     74% Matches
                                                273 Mismatches
                                                                       81
Gaps
                      12 Conservative Substitutions
                                                                        0
                                                             10
                                                                       20
```

GTCAAACTGCAGCAGTCTGGGG \$81 1 \$1111111111111

```
30
                          50
                                   60
                                            70
                                                     80
   CAGAGCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAA--AGACAC
   CTGAGCTGGTAAAGCCTGGGGCTTCAGTGAAGATGTCCTGCAAGGCTTCTGGATAC-ACATTTACTAGCTAT
               150
                        160
                                 170
                                          180
       100
                110
                         120
                                  130
                                           140
                                                    150
                                                             160
   CTATATGCACTGGGTGAAGCAGAGGCCTGAACAGGGCCTGGAGTGGATTGGAAGGATTGATCCTGCGAGTGG
    GT-TATGCACTGGGTGAAGCAGAAGCCTGGGCAGGGCCTTGAGTGGATTGGATATATTAATCCTTACAATGA
       210
                220
                        230
                                 240
                                          250
                                                   260
                                                            270
      170
               180
                        190
                                200
                                         210
                                                  220
                                                           230
   CGATACTAAATATGACCCGAAGTTCCAGGTCAAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTG
                       1 [ ] ] ] ] ] ] ]
   TGGTACTAGTTACAATGAGCCCTTCAAAGGCAAGGCCACACTGACTTCAGACAAATCCTCCAGCACAGCCTA
     280
              290
                       300
                               310
                                        320
                                                 330
                                                          340
    240
             250
                      260
                               270
                                       280
                                                 290
                                                             300
   GCTGCAGCTCAGCAGCCTGACATCTGAGGACACTGCCGTCTACTACTGTGC-AGACGGA---ATGTGGGTAT
     CATGGAGCTCAGCAGCCTGACCTCTGAGGACTCTGCGGTCTATTACTGTGCAAGGAGGATCTACTTTGATTA
   350
            360
                     370
                              380
                                      390
                                               400
                                                        410
      310
               320
                        330
                                340
                                         350
   CAACGGGATATGCTCTGGACTTCTGGGGCCAAGGGACCACGGTCACCGTCTCCTCA
          CTCC----TATGTTATGGACTACTGGGGTCAAGGAACCTCGGTCACCGTCTCCTCA
 420
              430
                       440
                               450
                                        460
                                                 470
4. US-07-977-702-1 (1-360)
  022736
              RSV19 VH.
ID
     022736 standard; DNA; 348 BP.
AC
     022736;
DT
     13-AUG-1992 (first entry)
DF
     RSV19 VH.
ΚW
     VH; VK; donor; antibody; CDR; epitope; NEW; REI;
KW
     fusion protein; F protein; ss.
05
     Mus nusculus.
FH
     Key
                   Location/Qualifiers
FT
     misc_feature
                   91..105
FT
     /*tag= a
FT
     /label= CDR1
FT
                   149..199
     misc_feature
FT
     /#tag= b
FT
     /label= CDR2
FT
     misc_feature
                   296..316
FT
     /*tag= c
FT
     /label= CDR3
FT
     primer_bind
                   2..23
FT
     /≱tag= d
FT
     /note= "corresponds to primer sequence used"
FT
     primer_bind
                   316..349
FT
     /*tag= e
FT
     /note= "corresponds to primer sequence used"
PN
     W09204381-A.
PD
     19-MAR-1992.
PF
     11-SEP-1991; G01554.
PR
     11-SEP-1990; GB-019812.
PA
     (SCOT-) SCOTGEN LTD.
PI
     Harris WJ, Tempest PR, Taylor G;
```

DR

WPI; 92-114306/14.

11U X

120

130

```
PT
     New altered antibodies with donor MAb binding specificity for RSV
PT

    for treatment and prevention of human respiratory syncutial

PT
     virus infection
PS
     Disclosure; Fig 1; 72pp; English.
CC
     The sequences of RSV19 VH and VK are represented in @22736 and @22737
CC
     respectively. The donor antibody RSV19 is directed against epitope
CC
     417-438 of the fusion protein (F). The CDRs were identified (see
CC
     features), then the murine CDRs transferred to human frameworks by
CC
     site-directed mutagenesis, using as DNA templates human framework
CC
     regions of the NEW (heavy) and REI (light) proteins.
CC
     The altered antibodies are used to prevent or treat RSV infections
CC
     in humans, e.g. for prevention they are given (each 6 weeks during
CC
     the RSV season) at 1-20 mg/kg parenterally or 0.2-2 mg/kg
CC
     intranasally. Since the antibodies are predominantly
CC
     human, they are unlikely to cause much immune response.
SQ
     Sequence
              348 BP;
                        84 A;
                                 91 C;
                                         95 G;
                                                 76 T;
SO
     2 Others;
Initial Score
                   256 Optimized Score =
                                           299
                                                Significance = 17.51
Residue Identity =
                   84% Matches
                                            305
                                                Mismatches
                                                                40
                    15 Conservative Substitutions
Gaps
                                                                 0
             10
                      20
                               30
                                        40
                                                 50
      GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCT
      CAGGTCCAGCTGCAGSAGTCWGGGACAGAGCTTGAGAGGTCAGGGGGCCTCAGTCAAGTTGTCCTGCACAGCT
          10
                   20
                            30
                                     40
                                              50
                                                       60
  70
           80
                    90
                            100
                                     110
                                              120
                                                       130
                                                                140
   TCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAGCAGGGCCTGAACAGGGCCTGGAGTGGATT
   TCTGGCTTCAACATTAAAGACTACTATATGCACTGGATGAAGCAGAGGCCTGACCAGGGCCTGGAGTGGATT
        80
                 90
                         100
                                  110
                                           120
                                                     130
                                                             140
        150
                          170
                 160
                                   180
                                            190
                                                     200
                                                              210
   GGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTCAAGGCCACTATTACAGCG
   GGATGGATTGATCCTGAGAATGATGATGTTCAATATGCCCCGAAGTTCCAGGGCAAGGCCACTATGACTGCA
      150
               160
                        170
                                 180
                                          190
                                                   200
                                                            210
       220
                230
                        240
                                  250
                                           260
                                                   270
                                                            280
   GACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGACACTGCCGTCTACTACTGT
   GACACGTCCTCCAACACAGCCTACCTGCAGCTCACCAGCCTGACATTTGAGGACACTGCCGTCTATTTCTGT
    220
             230
                      240
                               250
                                        260
                                                 270
                                                          280
     290
              300
                       310
                                320
                                         330
                                                  340
                                                           350
   GCAGACGGAATGTGGGTATCAACGGGATATGCTCTGGACTTCTGGGGCCAAGGGACCACGGTCACCGTCTCC
          \Pi\Pi
                   111 - 111
                            - 11
                                  -----AAT-----TCATGGGGGAGTGACTTTGACCACTGGGGCCAAGGGACCACGGTCACCGTCTCC
         290
                        300
                                  310
                                           320
                                                   330
                                                            340
   360
   TCA
   \Pi\Pi
   TCA
     X
```

5. US-07-977-702-1 (1-360) 028739 cDNA of VH425 antibody cloned into pUC18.

ID @28739 standard; DNA; 501 BP.

AC @28739;

DT 01-MAR-1993 (first entry)

P-P5UB: K248U/.

```
DΈ
      CDNA OF VH425 antibody cloned into pUC18.
 KW
      Monoclonal antibody; complementarity determining region; framework;
 KW
      antigens; tumour; melanoma; carcinoma; glioma; variable; heavy;
 K₩
      light; chain; ss.
 OS
      Synthetic.
 FH
     Key
                     Location/Qualifiers
 FT
      CDS
                     30..450
 FT
      /*tag= a
 FT
      sig_peptide
                     .30..87
 FT
      /#tag= b
 FT
     mat_peptide
                     88..450
 FT
      /#tag= c
 FT
      primer_bind
                     10..32
 FT
      /*tag= d
 FT
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 FT
      primer_bind
                     12..32
 FT
      /#tag= e
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      /note= "PCR primer binding site"
 FT
      primer_bind
                     436..465
 FT
     /*tag= f
      /note= "3'-5' PCR primer binding site"
 FT
PN
     W09215683-A.
PD
     17-SEP-1992.
PF
     04-MAR-1992; E00480.
 PR
     06-MAR-1991; EP-103389.
PA
      (MERE ) MERCK PATENT GMBH.
PI
     Bendig MM, Kettleborough CA, Saldanha J;
 DR
     WPI; 92-331729/40.
 DR
     P-PSBD; R27049.
 PT
     Human monoclonal antibodies binding to human receptors - for
 PT
     treatment and diagnosis of tumours, e.g. melanoma and carcinoma
 PS
     Disclosure; Fig 2; 89pp; English.
 CC
     The cDNA sequence encoding the variable heavy chain of monoclonal
 CC
      antibody 425 was prepd. synthetically and mutations made to the 5'
 CC
      and 3' ends to allow for cloning into HCMV expression vectors.
 CC
     Donor splice sites were recreated in the 3' flanking regions to
 CC
      allow correct splicing of the variable and constant regions. The
 CC
     5' sequence was altered to introduce an initiation codon. These
 CC
     mutations were carried out using the PCR primers shown in the
 CC
     features table. The cloned chimeric antibody contg. 425 VL and
 CC
     425 VH regions were cotransfected into COS cells to confirm cloning
 CC
     of the correct mouse base variable sequence. The cloned antibody
 CC
     may be used int the prodn.of reshaped or humanised antibodies which
 CC
      are less immunogenic than native antibodies and may be used to combat
 CC
      e.g. glioma, melanoma or carcinoma.
 CC
      See also @27040-1.
 SQ
      Sequence
                501 BP;
                           127 A;
                                     146 C;
                                               124 G;
                                                         104 T;
Initial Score
                     256 Optimized Score =
                                                269 Significance = 17.51
Residue Identity =
                     75% Natches
                                                275 Mismatches
                                                                       82
Gaps
                       6 Conservative Substitutions
                                                                        0
                                                             10
                                                                       20
                                                     GTCAAACTGCAGCAGTCTGGGG
                                                     111 | 11111 || 111111
    GCTATATCATCCTCTTTTTGGTAGCAACAGCTACAGATGTCCACTCCCAGGTCCAGCTGCAACAACCTGGGG
            50
                     60
                               70
                                         80
                                                   90
                                                            100
                                                                      110
          30
                   40
                             50
                                       60
                                                 70
                                                           80
    CAGAGCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTA-AAGACACC
    CTGAACTGGTGAAGCCTGGGGCTTCAGTGAAGTTGTCCTGCAAGGCTTCCGGCTACACCTTCACCAGCCA-C
         120
                  130
                            140
                                      150
                                                160
                                                          170
                                                                    180
        100
                 110
                           120
                                     130
                                               140
                                                         150
```

TATATGCACTGGGTGAAGCAGAGGCCTGAACAGGGCCTGGAGTGGATTGGAAGGATTGATCCTGCGAGTGGC

```
TGGATGCACTGGGTGAAGCAGAGGGCTGGACAAGGCCTTGAGTGGATCGGAGAGTTTAATCCCAGCAACGGC
       190
                200
                         210
                                 220
                                          230
                                                   240
                                                            250
     170
              180
                       190
                                200
                                        210
                                                 220
                                                          230
   GATACTAAATATGACCCGAAGTTCCAGGTCAAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGG
     260
              270
                       280
                               290
                                        300
                                                 310
                                                          320
   240
            250
                     260
                              270
                                       280
                                                290
                                                          300
   CTGCAGCTCAGCAGCCTGACATCTGAGGACACTGCCGTCTACTACTGTG-CAGAC-GGAATGTGGGTATCAA
    ATGCAACTCAGCAGCCTGACATCTGAGGACTCTGCGGTCTATTACTGTGCCAGTCGGGACTATGATTA-CGA
   330
            340
                     350
                              360
                                       370
                                                380
                                                        390
   310
            320
                     330
                              340
                                       350
                                                 X
   CGGGATATGCTCTGGACTTCTGGGGCCAAGGGACCACGGTCACCGTCTCCTCA
          CGGACGGTACT-TTGACTACTGGGGCCAAGGCACCACTCTCACAGTCTCCTCAGCCAAAACAACACCCCATC
  400
            410
                     420
                              430
                                       440
                                               450
   GGTCTATCCACTGGATTCCTCTAGAGTCGAC
 470
          480
                   490
                            500
6. US-07-977-702-1 (1-360)
  @23863
              ScFvB18 construct mutant #6.
ID
     023863 standard; DNA; 770 BP.
AC
     023863;
DT
     21-MAY-1992 (first entry)
DE
     ScFvB18 construct mutant #6.
KW
     Fd; bacteriophage; gene III; filamentous; phagemid; capsid; coat;
K₩
     pilus; g3p; binding; adsorption; gene VIII; diverse repertoire;
KW
     specific binding pairs; replicable genetic display package; ds.
08
     Synthetic.
FH
     Key
                   Location/Qualifiers
FT
     CDS
                   1..770
FT
     /*taq= a
FT
     /product= scFvB18
FT
     mutation
                   734
FT
     /#tag= b
FT
     /note= "c -> t; Thr -> Ile (VL FR4)"
PN
     WD9201047-A.
PD
     23-JAN-1992.
PF
     10-JUL-1991; G01134.
PR
     10-JUL-1990; GB-015198.
PR
     19-DCT-1990; GB-022845.
PR
     12-NOV-1990; GB-024503.
PR
     06-MAR-1991; GB-004744.
PR
     15-MAY-1991; GB-010549.
PA
     (CAMB-) CAMBRIDGE ANTIBODY.
     (MEDI-) MED RES COUNCIL.
PA
PΙ
     McCafferty J. Pope AR, Johnson KS, Hoogenboom HRJ, Griffiths AD;
PΙ
     Jackson RH, Holliger KP, Marks JD, Clackson TP, Chiswell DJ;
PI
     Winter GP, Bonnert TP;
DR
     ₩PI; 92-056862/07.
DR
     P-PSDB; R22587.
PT
     Producing members of specific binding pairs - by expression in
PT
     recombinant host cells with a secreting replicable genetic
PT
     display package.
PS
     Example 38; Fig 44; 209pp; English.
CC
     The sequence encodes an antibody scFv fragment directed against 4-
CC
     hydroxy-3-nitrophenylacetic acid (NP). HV and LV chains of an MAb
```

against NP were separately amplified and reassembled to form the

CC

```
conscruct, which was then ligated into the 10 deue III contd. Aec-
 CC
     tor, fdCAT2, derived from fdTPs/Xh.(See @21095). The clone having
 CC
     the scFvB18 sequence (see G21100) fused in frame to gene III was
 CC
     designated fdCAT2scFvB18. Alternatively the fragment was cloned
 CC
     into fdDDGKan (fdCAT2 with the tet gene replaced by a kanamycin
 CC
     resistance gene) to give fdDOGKanscFvB18, or into the phagenid
 CC
     pHEN1 to create pHEN1-scFvB18. The constructs were used to test
 CC
     the effect of using mutator strains to increase the diversity of
 CC
     the cloned genes. The strains NR9046mutD5: NR9046 mutD5::Tn10 and
 CC
     NR9046mutT1: NR9046mutT1::Tn10 were constructed by P1 transduction.
 CC
     After 4 rounds of mutation and screening, 40 phage inserts were
 CC
     sequenced. They each displayed single mutations in 6 different
 CC
     positions, five being in the light chain region. More than 70% of
 CC
     the mutations occurred at positions 724 and 725 changing the first
 CC
     Gly in the J segment (framework 4) to Ser (in 21 cases here) or
 CC
     Asp (in 3 cases). The mutant shown here (see feature table for
 CC
     details of the mutation) occurred once. The mutant fragments had
 CC
     affinities for NP which were comparable to the wild-type scFv
 CC
     fragment (20nM).
 CC
     N.B. fdCAT2 is also referred to as fd-tet-DOG1 and fdDOG1.
 CC
     See also 021092-100, 103-116, 126-131; 023463, 465-495, 693-719,
 CC
     736-738, and 793-862.
 SO
     Sequence
               770 BP;
                          188 A;
                                   196 C;
                                             214 G;
                                                      172 T;
                    236 Optimized Score =
Initial Score
                                              280 Significance = 16.05
Residue Identity =
                                              288 Mismatches
                    79% Matches
Gaps
                     11 Conservative Substitutions
                                                                    0
                               10
                                        20
                                                 30
                                                           40
                                                                    50
                       GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTC
                       TTCTATTCTCACAGTGCACAGGTCCAGCTGCAGCAGTCTGGGGGCTTGTGAAGCCTGGGGCTTCAGTG
           10
                    20 X
                              30
                                       40
                                                50
                                                          60
                                                                   70
          60
                   70
                             80
                                      90
                                               100
                                                        110
                                                                  120
   AAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAGCAGAGGCCTGAA
    1 11 1111111111111111111111111111111
   AAGCTGTCCTGCAAGGCTTCTGGCTACACCTTCACCAGCTACTGGATGCACTGGGTGAAGCAGAGGCCTGGA
         80
                  90
                                    110
                           100
                                              120
                                                       130
                                                                140
       130
                 140
                          150
                                   160
                                             170
                                                      180
                                                                190
   CAGGGCCTGGAGTGGAATTGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTC
      CGAGGCCTTGAGTGGATTGGAAGGATTGATCCTAATAGTGGTGGTACTAAGTACAATGAGAAGTTCAAGAGC
      150
               160
                         170
                                  180
                                            190
                                                     200
                                                               210
     200
               210
                        220
                                 230
                                           240
                                                    250
                                                              260
   AAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGAC
    AAGGCCACACTGACTGTAGACAAACCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGACATCTGAGGAC
    220
             230
                       240
                                250
                                          260
                                                   270
                                                             280
   270
             280
                       290
                                 300
                                             310
                                                      320
   ACTGCCGTCTACTACTGTGC-AGA-CGGAATGTGG--GTATCAACGGGATATGCTCTGGACTTCTGGGGCCA
    1111 11111 11 11111 111
                             11 1 11 111 1 11
                                                 TCTGCGGTCTATTATTGTGCAAGATACGACTACGGTAGTAGCTAC---TA--CT-TTGACTACTGGGGCCA
  290
            300
                     310
                               320
                                        330
                                                        340
                                                                 350
     340
               350
                          X
   AGGGACCACGGTCACCGTCTCCTCA
    11111111111111111111111111
   AGGGACCACGGTCACCGTCTCCTCAGGTGGAGGCGGTTCAGGCGGAGGTGGCTCTGGCGGTGGCGGATCCCA
       360
                370
                          380
                                   390
                                             400
                                                      410
                                                                420
```

...

```
7. US-07-977-702-1 (1-360)
   023862
                ScFvB18 construct mutant #5.
ID
      Q23862 standard; DNA; 770 BP.
AC
      023862;
DT
     21-MAY-1992 (first entru)
     ScFvB18 construct autant #5.
DE
KW
     Fd; bacteriophage; gene III; filamentous; phagemid; capsid; coat;
KW
      pilus; g3p; binding; adsorption; gene VIII; diverse repertoire;
KW
     specific binding pairs; replicable genetic display package; ds.
OS
      Synthetic.
FH
     Key
                      Location/Qualifiers
FT
     CDS
                      1..770
FT
      /*tag= a
FT
     /product= scFvB18
FT
      mutation
                      725
FT
     /*tag= b
FT
     /note= "g -> a; Gly -> Asp (VL FR4)"
PN
     W09201047-A.
PD
     23-JAN-1992.
PF
     10-JUL-1991; G01134.
PR
     10-JUL-1990; GB-015198.
PR
     19-0CT-1990; GB-022845.
PR
     12-NOV-1990; GB-024503.
     06-MAR-1991; GB-004744.
PR
     15-MAY-1991; GB-010549.
PA
     (CAMB-) CAMBRIDGE ANTIBODY.
PA
     (MEDI-) MED RES COUNCIL.
PΙ
     McCafferty J. Pope AR. Johnson KS. Hoogenboom HRJ. Griffiths AD;
PΙ
      Jackson RH, Holliger KP, Marks JD, Clackson TP, Chiswell DJ;
PΙ
     Winter GP, Bonnert TP;
DR
     WPI; 92-056862/07.
DR
     P-PSDB; R22586.
PT
     Producing members of specific binding pairs - by expression in
PT
     recombinant host cells with a secreting replicable genetic
     display package.
PT
PS
      Example 38; Fig 44; 209pp; English.
CC
     The sequence encodes an antibody scFv fragment directed against 4-
      hydroxy-3-nitrophenylacetic acid (NP). HV and LV chains of an MAb
CC
CC
      against NP were separately amplified and reassembled to form the
CC
     construct, which was then ligated into the fd gene III contq. vec-
CC
     tor, fdCAT2, derived from fdTPs/Xh. (See @21095). The clone having
CC
      the scFvB18 sequence (see @21100) fused in frame to gene III was
CC
      designated fdCAT2scFvB18. Alternatively the fragment was cloned
CC
      into fdDOGKan (fdCAT2 with the tet gene replaced by a kanamycin
CC
      resistance gene) to give fdDOGKanscFvB18, or into the phagemid
CC
      pHEN1 to create pHEN1-scFvB18. The constructs were used to test
      the effect of using mutator strains to increase the diversity of
CC
CC
      the cloned genes. The strains NR9046mutD5: NR9046 mutD5::Tn10 and
CC
      NR9046mutT1: NR9046mutT1::Tn10 were constructed by P1 transduction.
CC
      After 4 rounds of mutation and screening, 40 phage inserts were
      sequenced. They each displayed single mutations in 6 different
CC
CC
     positions, five being in the light chain region. More than 70% of
CC
     the mutations occurred at positions 724 and 725 changing the first
CC
     Gly in the J segment (framework 4) to Ser (in 21 cases), or Asp (in
CC
     3 cases as shown here). The mutant fragments had affinities for
CC
     NP which were comparable to the wild-type scFv fragment (20nM).
CC
     N.B. fdCAT2 is also referred to as fd-tet-DOG1 and fdDOG1.
CC
     See also @21092-100, 103-116, 126-131; @23463, 465-495, 693-719,
CC
     736-738, and 793-863.
SQ
     Sequence
                770 BP;
                            189 A;
                                      197 C;
                                                213 G;
                                                          171 T;
Initial Score
                      236 Optimized Score =
                                                 280 Significance = 16.05
Residue Identity =
                      79% Matches
                                                 288 Mismatches
                                            =
```

```
10
                                      20
                                               30
                                                                  50
                      GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTC
                      TTCTATTCTCACAGTGCACAGGTCCAGCTGCAGCAGTCTGGGGCTGAGCTTGTGAAGCCTGGGGCTTCAGTG
                   20 X
          10
                            30
                                     40
                                              50
                                                       60
         60
                   70
                           80
                                     90
                                             100
                                                      110
                                                               120
   AAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAGCAGAGGCCTGAA
                                     AAGCTGTCCTGCAAGGCTTCTGGCTACACCTTCACCAGCTACTGGATGCACTGGGTGAAGCAGAGGCCTGGA
         80
                  90
                          100
                                   110
                                            120
                                                     130
                                                              140
       130
                140
                         150
                                           170
                                  160
                                                    180
                                                             190
   CAGGGCCTGGAGTGGAATTGGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTC
      11111111111
   CGAGGCCTTGAGTGGATTGGAAGGATTGATCCTAATAGTGGTGGTACTAAGTACAATGAGAAGTTCAAGAGC
      150
                                          190
               160
                        170
                                 180
                                                   200
     200
              210
                       220
                                         240
                                230
                                                  250
                                                           260
   AAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGAC
   1411111 | 11 | 1141 | 11111 | 111111
                                          31111111111111111111111111111111
   AAGGCCACACTGACTGTAGACAAACCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGACATCTGAGGAC
    220
                      240
             230
                               250
                                        260
                                                 270
                                                          280
   270
                      290
            280
                                300
                                           310
                                                    320
                                                             330
   ACTGCCGTCTACTACTGTGC-AGA-CGGAATGTGG--GTATCAACGGGATATGCTCTGGACTTCTGGGGCCA
    TCTGCGGTCTATTATTGTGCAAGATACGACTACGGTAGTAGCTAC----TA--CT-TTGACTACTGGGGCCA
  290
           300
                    310
                             320
                                      330
                                                      340
                                                               350
     340
              350
                         X
   AGGGACCACGGTCACCGTCTCCTCA
   AGGGACCACGGTCACCGTCTCCTCAGGTGGAGGCGGTTCAGGCGGAGGTGGCTCTGGCGGTGGCGGATCCCA
       360
                370
                         380
                                  390
                                           400
                                                    410
                                                             420
   GGC
8. US-07-977-702-1 (1-360)
  023861
              ScFvB18 construct mutant #4.
ID
     023861 standard; DNA; 770 BP.
AC
     023861;
DT
     21-MAY-1992 (first entru)
DE
     ScFvB18 construct mutant #4.
     Fd; bacteriophage; gene III; filamentous; phagemid; capsid; coat;
KH
KW
     pilus; g3p; binding; adsorption; gene VIII; diverse repertoire;
KW
     specific binding pairs; replicable genetic display package; ds.
05
     Synthetic.
FH
     Key
                   Location/Qualifiers
FT
     CDS
                    1..770
FT
     /#tag= a
FT
     /product= scFvB18
FT
     mutation
                   724
FT
     /*tag= b
FT
     /note= "g -> a; Gly -> Ser (VL FR4)"
PN
     W09201047-A.
PD
     23-JAN-1992.
PF
     10-JUL-1991; G01134.
PR
     10-JUL-1990; GB-015198.
PR
     19-0CT-1990; GB-022845.
```

PR

12-NOV-1990; GB-024503.

conservative bubstitutions

```
PR
      15-MAY-1991; GB-010549.
 PA
      (CAMB-) CAMBRIDGE ANTIBODY.
 PA
      (MEDI-) MED RES COUNCIL.
 PI
     McCafferty J. Pope AR. Johnson KS. Hoogenboom HRJ. Griffiths AD:
 PI
      Jackson RH, Holliger KP, Marks JD, Clackson TP, Chiswell DJ;
 PI
     Winter GP, Bonnert TP;
 DR
     WPI; 92-056862/07.
 DR
     P-PSDB; R22585.
 PT
     Producing members of specific binding pairs - by expression in
 PT
     recombinant host cells with a secreting replicable genetic
 PT
     display package.
 PS
     Example 38; Fig 44; 209pp; English.
 CC
     The sequence encodes an antibody scFv fragment directed against 4-
     hydroxy-3-nitrophenylacetic acid (NP). HV and LV chains of an MAb
 CC
 CC
     against NP were separately amplified and reassembled to form the
 CC
     construct, which was then ligated into the fd gene III contg. vec-
 CC
     tor, fdCAT2, derived from fdTPs/Xh.(See 021095). The clone having
 CC
     the scFvB18 sequence (see @21100) fused in frame to gene III was
 CC
     designated fdCAT2scFvB18. Alternatively the fragment was cloned
CC
     into fdDOGKan (fdCAT2 with the tet gene replaced by a kanamycin
 CC
     resistance gene) to give fdDOGKanscFvB18, or into the phagemid
CC
     pHEN1 to create pHEN1-scFvB18. The constructs were used to test
CC
     the effect of using mutator strains to increase the diversity of
CC
     the cloned genes. The strains NR9046mutD5: NR9046 mutD5::Tn10 and
CC
     NR9046mutT1: NR9046mutT1::Tn10 were constructed by P1 transduction.
CC
     After 4 rounds of mutation and screening, 40 phage inserts were
CC
     sequenced. They each displayed single mutations in 6 different
CC
     positions, five being in the light chain region. More than 70% of
CC
     the mutations occurred at positions 724 and 725 changing the first
 CC
     Gly in the J segment (framework 4) to Ser (in 21 cases, as shown
CC
     here) or Asp (in 3 cases). The mutant fragments had affinities
CC
     for NP which were comparable to the wild-type scFv fragment (20nM).
CC
     N.B. fdCAT2 is also referred to as fd-tet-DDG1 and fdDDG1.
CC
     See also 021092-100, 103-116, 126-131; 023463, 465-495, 693-719,
     736-738, and 793-863.
CC
50
     Sequence
                770 BP;
                          189 A;
                                    197 C;
                                              213 G;
                                                       171 T;
Initial Score
                     236 Optimized Score =
                                               280 Significance = 16.05
Residue Identity =
                     79% Matches
                                               288 Mismatches
Gaps
                     11 Conservative Substitutions
                                                                      0
                               10
                                         20
                                                   30
                                                            40
                       GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTC
                        TTCTATTCTCACAGTGCACAGGTCCAGCTGCAGCAGTCTGGGGGCTTGTGAAGCCTGGGGGCTTCAGTG
           10
                     20 X
                              30
                                        40
                                                 50
                                                           60
                                                                     70
          60
                   70
                             80
                                       90
                                               100
                                                         110
                                                                   120
   AAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAGCAGAGGCCTGAA
    AAGCTGTCCTGCAAGGCTTCTGGCTACACCTTCACCAGCTACTGGATGCACTGGGTGAAGCAGAGGCCTGGA
         80
                   90
                           100
                                     110
                                               120
                                                        130
                                                                  140
       130
                 140
                          150
                                    160
                                              170
                                                       180
                                                                 190
   CAGGGCCTGGAGTGGAATGGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTC
      CGAGGCCTTGAGTGGATTGGAAGGATTGATCCTAATAGTGGTGGTACTAAGTACAATGAGAAGTTCAAGAGC
      150
                160
                         170
                                   180
                                            190
                                                      200
                                                                210
     200
               210
                        220
                                  230
                                            240
                                                     250
                                                               260
   AAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGAC
   1111111 | 11 | 1111
                         111111 1111111
                                            11111111111111111111111111111111111
   AAGGCCACACTGACTGTAGACAAACCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGACATCTGAGGAC
    220
              230
                       240
                                 250
                                           260
                                                    270
```

U6-MAK-1771; G5-UU4/44.

```
ACTGCCGTCTACTACTGTGC-AGA-CGGAATGTGG--GTATCAACGGGATATGCTCTGGACTTCTGGGGCCA
     11 11 1 1111 111111111
    TCTGCGGTCTATTATTGTGCAAGATACGACTACGGTAGTAGCTAC----TA--CT-TTGACTACTGGGGCCA
   290
             300
                       310
                                320
                                          330
                                                           340
                                                                     350
      340
                350
    AGGGACCACGGTCACCGTCTCCTCA
    1111111111111111111111111
    AGGGACCACGGTCACCGTCTCCTCAGGTGGAGGCGGTTCAGGCGGAGGTGGCTCTGGCGGTGGCGGATCCCA
                  370
                                     390
                                               400
                                                          410
   GGC
9. US-07-977-702-1 (1-360)
   023860
               ScFvB18 construct mutant #3.
      Q23860 standard; DNA; 770 BP.
AC
      023860;
     21-MAY-1992 (first entry)
DT
DE
      ScFvB18 construct autant #3.
     Fd; bacteriophage; gene III; filamentous; phagemid; capsid; coat;
K₩
KW
      pilus; g3p; binding; adsorption; gene VIII; diverse repertoire;
KW
      specific binding pairs; replicable genetic display package; ds.
OS
      Synthetic.
FH
      Key
                     Location/Qualifiers
FT
      CDS
                     1..770
      /*tag= a
FT
     /product= scFvB18
FT
      Mutation
                     706
FT
      /*tag= b
      /note= "a -> g; Ser -> Gly (VL CDR3)"
PN
     W09201047-A.
     23-JAN-1992.
      10-JUL-1991; G01134.
PR
      10-JUL-1990; GB-015198.
PR
      19-OCT-1990; GB-022845.
PR
      12-NOV-1990; GB-024503.
PR
     06-MAR-1991; GB-004744.
PR
     15-MAY-1991; GB-010549.
PA
      (CAMB-) CAMBRIDGE ANTIBODY.
PA
      (MEDI-) MED RES COUNCIL.
PΙ
      McCafferty J, Pope AR, Johnson KS, Hoogenboom HRJ, Griffiths AD;
      Jackson RH, Holliger KP, Marks JD, Clackson TP, Chiswell DJ;
PΙ
PI
      Winter GP, Bonnert TP;
DR
     WPI; 92-056862/07.
DR
     P-PSDB; R22584.
PT
      Producing members of specific binding pairs - by expression in
      recombinant host cells with a secreting replicable genetic
PT
      display package.
PS
      Example 38; Fig 44; 209pp; English.
CC
      The sequence encodes an antibody scFv fragment directed against 4-
CC
      hydroxy-3-nitrophenylacetic acid (NP). HV and LV chains of an MAb
CC
      against NP were separately amplified and reassembled to form the
CC
      construct, which was then ligated into the fd gene III contg. vec-
CC
      tor, fdCAT2, derived from fdTPs/Xh.(See @21095). The clone having
CC
      the scFvB18 sequence (see 021100) fused in frame to gene III was
CC
      designated fdCAT2scFvB18. Alternatively the fragment was cloned
CC
      into fdDOGKan (fdCAT2 with the tet gene replaced by a kanamycin
CC
      resistance gene) to give fdDOGKanscFvB18, or into the phagemid
CC
      pHEN1 to create pHEN1-scFvB18. The constructs were used to test
CC
      the effect of using mutator strains to increase the diversity of
      the cloned genes. The strains NR9046mutD5: NR9046 mutD5::Tn10 and
      NR9046mutT1: NR9046mutT1::Tn10 were constructed by P1 transduction.
```

ID

FT

FT

PD

PF

PT

CC

CC

300

310

```
CC
     sequenced. They each displayed single mutations in 6 different
 CC
     positions, five being in the light chain region. More than 70% of
 CC
     the mutations occurred at positions 724 and 725 changing the first
 CC
     Gly in the J segment (framework 4) to Ser (in 21 cases) or Asp (in
 CC
     3 cases). The mutant shown here (see feature table for details of
 CC
     mutation) occurred once. The mutant fragments had affinities for
 CC
     NP which were comparable to the wild-type scFv fragment (20nM).
 CC
     N.B. fdCAT2 is also referred to as fd-tet-DOG1 and fdDOG1.
 CC
     See also @21092-100, 103-116, 126-131; @23463, 465-495, 693-719,
 CC
     736-738, and 793-863.
 Se
     Sequence
              770 BP;
                          187 A;
                                   197 C;
                                             215 G;
                                                       171 T;
Initial Score
                    236 Optimized Score =
                                              280 Significance = 16.05
Residue Identity =
                     79% Matches
                                              288 Mismatches
Gaps
                     11 Conservative Substitutions
                                                                     0
                               10
                                        20
                                                  30
                                                           40
                                                                     50
                       GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTC
                       TTCTATTCTCACAGTGCACAGGTCCAGCTGCAGCAGTCTGGGGCTTGTGAAGCCTGGGGCTTCAGTG
           10
                    20 X
                              30
                                                 50
                                       40
                                                          60
                                                                    70
          60
                   70
                                      90
                             80
                                               100
                                                        110
                                                                  120
   AAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAGCAGAGAGCCTGAA
   111 11111111 1111111111 11 1 1 1
                                       AAGCTGTCCTGCAAGGCTTCTGGCTACACCTTCACCAGCTACTGGATGCACTGGGTGAAGCAGAGGCCTGGA
         80
                  90
                           100
                                    110
                                              120
                                                       130
                                                                 140
       130
                 140
                          150
                                    160
                                             170
                                                       180
                                                                190
   CAGGGCCTGGAGTGGAATTGGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTC
      11111 111111111111111111111111
                                     CGAGGCCTTGAGTGGATTGGAAGGATTGATCCTAATAGTGGTGGTACTAAGTACAATGAGAAGTTCAAGAGC
      150
                160
                         170
                                                               210
                                   180
                                            190
                                                      200
     200
               210
                        220
                                  230
                                           240
                                                     250
                                                              260
   AAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGAC
    11111111 | 11 | 1111
                           AAGGCCACACTGACTGTAGACAAACCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGACATCTGAGGAC
    220
              230
                       240
                                 250
                                          260
                                                   270
                                                             280
   270
             280
                       290
                                  300
                                             310
                                                      320
                                                                330
   ACTGCCGTCTACTACTGTGC-AGA-CGGAATGTGG--GTATCAACGGGATATGCTCTGGACTTCTGGGGCCA
    1111 11111 11 11111 111
                             11 | 11 |11 | 11
                                                  11 11 1 1111 111111111
   TCTGCGGTCTATTATTGTGCAAGATACGACTACGGTAGTAGCTAC----TA--CT-TTGACTACTGGGGCCA
  290
            300
                     310
                               320
                                        330
                                                        340
                                                                  350
     340
               350
   AGGGACCACGGTCACCGTCTCCTCA
   AGGGACCACGGTCACCGTCTCCTCAGGTGGAGGCGGTTCAGGCGGAGGTGGCTCTGGCCGGTGGCCGATCCCA
       360
                 370
                          380
                                   390
                                             400
                                                       410
                                                                420
   GGC
10. US-07-977-702-1 (1-360)
   023859
                ScFvB18 construct mutant #2.
ID
     023859 standard; DNA; 770 BP.
AC
     023859;
     21-MAY-1992 (first entry)
DT
```

After 4 rounds of mutation and screening, 40 phage inserts were

KW Fd; bacteriophage; gene III; filamentous; phagemid; capsid; coat;
KW pilus; g3p; binding; adsorption; gene VIII; diverse repertoire;

ScFvB18 construct mutant #2.

DE

```
specific binding pairs; replicable genetic display package; ds.
 05
      Synthetic.
 FH
      Key
                      Location/Qualifiers
 FT
      CDS
                      1..770
 FT
      /*tag= a
 FT
      /product= scFvB18
 FT
      mutation
 FT
      /*tag= b
 FT
      /note= "t -> g; Tyr -> Asp (VL CDR3)"
 PN
      WD9201047-A.
 PD
      23-JAN-1992.
 PF
      10-JUL-1991; G01134.
 PR
      10-JUL-1990; GB-015198.
 PR
      19-0CT-1990; GB-022845.
 PR
      12-NOV-1990; GB-024503.
 PR
      06-MAR-1991; GB-004744.
 PR
      15-MAY-1991; GB-010549.
 PA
      (CAMB-) CAMBRIDGE ANTIBODY.
 PA
      (MEDI-) MED RES COUNCIL.
 PΙ
      McCafferty J. Pope AR. Johnson KS. Hoogenboom HRJ, Griffiths AD;
 PΙ
      Jackson RH, Holliger KP, Marks JD, Clackson TP, Chiswell DJ;
 PΙ
      Winter GP, Bonnert TP;
 DR
      WPI; 92-056862/07.
      P-PSDB; R22583.
 DR
 PT
      Producing members of specific binding pairs - by expression in
 PT
      recombinant host cells with a secreting replicable genetic
 PT
      display package.
PS
      Example 38; Fig 44; 209pp; English.
 CC
      The sequence encodes an antibody scFv fragment directed against 4-
 CC
      hydroxy-3-nitrophenylacetic acid (NP). HV and LV chains of an MAb
 CC
      against NP were separately amplified and reassembled to form the
 CC
      construct, which was then ligated into the fd gene III contg. vec-
 CC
      tor, fdCAT2, derived from fdTPs/Xh.(See @21095). The clone having
 CC
      the scFvB18 sequence (see Q21100) fused in frame to gene III was
 CC
      designated fdCAT2scFvB18. Alternatively the fragment was cloned
 CC
      into fdDOGKan (fdCAT2 with the tet gene replaced by a kanamycin
 CC
      resistance gene) to give fdDOGKanscFvB18, or into the phagemid
 CC
      pHEN1 to create pHEN1-scFvB18. The constructs were used to test
CC
      the effect of using mutator strains to increase the diversity of
 CC
      the cloned genes. The strains NR9046mutD5: NR9046 mutD5::Tn10 and
 CC
      NR9046mutT1: NR9046mutT1::Tn10 were constructed by P1 transduction.
 CC
      After 4 rounds of mutation and screening, 40 phage inserts were
 CC
      sequenced. They each displayed single mutations in 6 different
 CC
      positions, five being in the light chain region. More than 70% of
 CC
      the mutations occurred at positions 724 and 725 changing the first
 CC
      Gly in the J segment (framework 4) to Ser (in 21 cases) or Asp (in
 CC
      3 cases). The mutant shown here (see feature table for details of
 CC
      mutation) occurred once. The mutation shown on the fig., of t \rightarrow c
 CC
      would result in the Tyr being replaced by His, contrary to the data
 CC
      in Table 9 of the specification which indicates that the substn. is
CC
      with Asp (i.e. t -> g). The mutant fragments had affinities for NP
CC
      which were comparable to the wild-type scFv fragment (20nM).
CC
      N.B. fdCAT2 is also referred to as fd-tet-DOG1 and fdDOG1.
 CC
      See also @21092-100, 103-116, 126-131; @23463, 465-495, 693-719,
CC
      736-738, and 793-863.
SQ
      Sequence
                770 BP;
                           188 A;
                                      197 C;
                                               215 G;
                                                          170 T;
Initial Score
                =
                      236 Optimized Score =
                                                 280 Significance = 16.05
Residue Identitu =
                      79% Matches
                                                 288 Mismatches
                                                                  =
                                                                        65
Gaps
                      11 Conservative Substitutions
                                                                   =
                                                                         0
                         X
                                 10
                                           20
                                                     30
                                                               40
                         GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTC
                         TTCTATTCTCACAGTGCACAGGTCCAGCTGCAGCAGTCTGGGGCTGAGCTTGTGAAGCCTGGGGCTTCAGTG
            10
                     20 X
                                30
                                          40
                                                   50
                                                              60
```

RΠ

```
60
                     70
                                80
                                          90
                                                   100
                                                             110
                                                                       120
    AAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAGCAGAGGCCTGAA
    111 111111111 1111111111 11 1 1 1
                                           AAGCTGTCCTGCAAGGCTTCTGGCTACACCTTCACCAGCTACTGGATGCACTGGGTGAAGCAGAGGCCTGGA
                    90
          80
                             100
                                        110
                                                  120
                                                            130
                                                                      140
        130
                  140
                            150
                                                 170
                                       160
                                                           180
                                                                     190
    CAGGGCCTGGAGTGGAATGGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTC
       11111 11111111111111111111111111
                                         11111 | 111111 | 1
    CGAGGCCTTGAGTGGATTGGAAGGATTGATCCTAATAGTGGTGGTACTAAGTACAATGAGAAGTTCAAGAGC
       150
                 160
                           170
                                      180
                                                190
                                                          200
      200
                210
                          220
                                    230
                                               240
                                                         250
                                                                   260
    AAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGAC
                             111111 11111111
    11111111 | 11 | 1111
                                                111111111111111111111111111111111111
    AAGGCCACACTGACTGTAGACAAACCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGACATCTGAGGAC
     220
               230
                         240
                                   250
                                              260
                                                        270
                                                                  280
    270
              280
                         290
                                    300
                                                 310
                                                           320
                                                                     330
    ACTGCCGTCTACTACTGTGC-AGA-CGGAATGTGG--GTATCAACGGGATATGCTCTGGACTTCTGGGGCCA
     1111 11111 11 11111 111
                               11 1 11 111 1 11
                                                      11 11 1 1111 111111111
    TCTGCGGTCTATTATTGTGCAAGATACGACTACGGTAGTAGCTAC----TA--CT-TTGACTACTGGGGCCA
   290
             300
                       310
                                 320
                                            330
                                                             340
                                                                       350
      340
                350
                            Y
    AGGGACCACGGTCACCGTCTCCTCA
    111111111111111111111111111
    AGGGACCACGGTCACCGTCTCCTCAGGTGGAGGCGGTTCAGGCGGAGGTGGCTCTGGCGGTGGCGGATCCCA
        360
                            380
                  370
                                      390
                                                 400
                                                           410
                                                                     420
    GGC
11. US-07-977-702-1 (1-360)
                 ScFvB18 construct.
 ID
      021100 standard; DNA; 770 BP.
 AC
      021100;
      21-MAY-1992 (first entry)
 DT
 DE
      ScFvB18 construct.
 KW
      Fd; bacteriophage; gene III; filamentous; phagemid; capsid; coat;
 K₩
      pilus; g3p; binding; adsorption; gene VIII; diverse repertoire;
 K₩
      specific binding pairs; replicable genetic display package; ds.
 05
      Synthetic.
FH
    Keu
                      Location/Qualifiers
FT
      CDS
                      1..770
 FT
      /*tag= a
FT
      /product= scFvB18
 FT
      mutation
                      308
FT
      /*tag= b
FT
      /note= "c -> t; Ala -> Val (VH FR3); @23858"
FT
      mutation
                      703
FT
      /*tag= c
      /note= "t -> g; Tyr -> Asp (VL CDR3); @23859"
FT
FT
      mutation
                      706
FT
      /*tag= d
FT
      /note= "a -> g; Ser -> Gly (VL CDR3); 023860"
FT
      mutation
                      724
FT
      /*tag= e
FT
      /note= "g -> a; Gly -> Ser (VL FR4); @23861"
FT
                      725
      mutation
FT
      /*tag= f
FT
      /note= "g -> a; Gly -> Asp (VL FR4); @23862"
```

FT

mutation

734

```
TΙ
 FT
      /note= "c -> t; Thr -> Ile (VL FR4); @23863"
 PN
     ₩09201047-A.
 PD
     23-JAN-1992.
 PF
      10-JUL-1991; G01134.
 PR
      10-JUL-1990; GB-015198.
 PR
      19-0CT-1990; GB-022845.
 PR
      12-NOV-1990; GB-024503.
 PR ·
     06-MAR-1991; GB-004744.
 PR
      15-MAY-1991; GB-010549.
 PA
      (CAMB-) CAMBRIDGE ANTIBODY.
 PA
      (MEDI-) MED RES COUNCIL.
 PΙ
     McCafferty J, Pope AR, Johnson KS, Hoogenboom HRJ, Griffiths AD;
 PΙ
     Jackson RH, Holliger KP, Marks JD, Clackson TP, Chiswell DJ;
 ΡI
     Winter GP, Bonnert TP;
     WPI; 92-056862/07.
 DR
 DR
     P-PSDB; R22568.
 PT
     Producing members of specific binding pairs - by expression in
 PT
     recombinant host cells with a secreting replicable genetic
 PT
     display package.
 PS
     Example 38; Fig 44; 209pp; English.
 CC
     The sequence encodes an antibody scFv fragment directed against 4-
 CC
     hydroxy-3-nitrophenylacetic acid (NP). HV and LV chains of an MAb
 CC
     against NP were separately amplified and reassembled to form the
 CC
     construct, which was then ligated into the fd gene III contg. vec-
 CC
     tor, fdCAT2, derived from fdTPs/Xh. (See @21095). The clone having
 CC
     the scFvB18 sequence fused in frame to gene III was designated
 CC
     fdCAT2scFvB18. Alternatively the fragment was cloned into fdDOGKan
 CC
      (fdCAT2 with the tet gene replaced by a kanamycin resistance gene)
 CC
     to give fdDOGKanscFvB18, or into the phagemid pHEN1 to create pHEN1-
 CC
     scFvB18. The constructs were used to test the effect of usingmuta-
 CC
     tor strains to increase the diversity of the cloned genes. The
 CC
     strains NR9046mutD5: NR9046 mutD5::Tn10 and NR9046mutT1: NR9046
 CC
     mutT1::Tn10 were constructed by P1 transduction. After 4 rounds of
 CC
     mutation and screening, 40 phage inserts were sequenced. They each
 CC
     displayed single mutations in 6 different positions, five being in
 CC
     the light chain region. More than 70% of the mutations occurred at
 CC
     positions 724 and 725 changing the first Gly in the J segment
 CC
      (framework 4) to Ser (in 21 cases) or Asp (in 3 cases). The
 CC
     mutant fragments had affinities for NP which were comparable to the
 CC
     wild-type scFv fragment (20nM).
 CC
     N.B. fdCAT2 is also referred to as fd-tet-DOG1 and fdDOG1.
 CC
     See also 021092-99, 103-116, 126-131; 023463, 465-495, 693-719,
CC
     736-738, and 793-863.
 SQ
     Sequence
                770 BP;
                          188 A;
                                    197 C;
                                              214 G;
                                                        171 T;
Initial Score
                     236 Optimized Score =
                                               280 Significance = 16.05
Residue Identity =
                     79% Matches
                                               288 Mismatches
                                                                     65
Gaps
                      11 Conservative Substitutions
                                                                      0
                               10
                                         20
                                                   30
                                                            40
                        GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTC
                        TTCTATTCTCACAGTGCACAGGTCCAGCTGCAGCAGTCTGGGGGCTGAGCTTGTGAAGCCTGGGGCTTCAGTG
           10
                     20 X
                              30
                                        40
                                                  50
                                                           60
                                                                     70
                    70
                                       90
          60
                             80
                                                100
                                                         110
   AAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAGCAGAGGCCTGAA
    AAGCTGTCCTGCAAGGCTTCTGGCTACACCTTCACCAGCTACTGGATGCACTGGGTGAAGCAGAGGCCTGGA
         80
                   90
                           100
                                     110
                                               120
                                                        130
                                                                  140
                 140
       130
                          150
                                              170
                                    160
                                                        180
                                                                 190
   CAGGGCCTGGAGTGGAATTGGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTC
```

CGAGGCCTTGAGTGGATTGGAAGGATTGATCCTAATAGTGGTGGTACTAAGTACAATGAGAAGTTCAAGAGC

```
200
                210
                         220
                                   230
                                             240
                                                       250
                                                                 260
    AAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGAC
    AAGGCCACACTGACTGTAGACAAACCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGACATCTGAGGAC
    220
              230
                        240
                                  250
                                            260
                                                      270
                                                                280
    270
             280
                        290
                                   300
                                               310
                                                         320
                                                                   330
    ACTGCCGTCTACTACTGTGC-AGA-CGGAATGTGG--GTATCAACGGGATATGCTCTGGACTTCTGGGGCCA
    1111 11111 11 11111 111
                              11 | 11 | 11 | 11
                                                    11 11 1 1111 11111111
    TCTGCGGTCTATTATTGTGCAAGATACGACTACGGTAGTAGCTAC----TA--CT-TTGACTACTGGGGCCA
   290
             300
                      310
                                320
                                          330
                                                           340
                                                                     350
      340
               350
    AGGGACCACGGTCACCGTCTCCTCA
    11111111111111111111111111
    AGGGACCACGGTCACCGTCTCCTCAGGTGGAGGCGGTTCAGGCGGAGGTGGCTCTGGCGGTGGCGATCCCA
        360
                  370
                           380
                                     390
                                               400
                                                         410
                                                                   420
   GGC
12. US-07-977-702-1 (1-360)
                ScFvB18 construct mutant #1.
    923858
 ID
      Q23858 standard; DNA; 770 BP.
 AC
      023858;
     21-MAY-1992 (first entry)
DT
     ScFvB18 construct mutant #1.
 KW
      Fd; bacteriophage; gene III; filamentous; phagemid; capsid; coat;
 KW
      pilus; g3p; binding; adsorption; gene VIII; diverse repertoire;
 K₩
      specific binding pairs; replicable genetic display package; ds.
 05
      Synthetic.
FH
      Key
                     Location/Qualifiers
FT
      CDS
                     1..770
FT
      /#tag= a
FT
      /product= scFvB18
FT
      mutation
                     308
FT
      /*tag= b
 FT
      /note= "c -> t; Ala -> Val (VH FR3)"
 PN
      W09201047-A.
PD
      23-JAN-1992.
 PF
      10-JUL-1991; G01134.
 PR
      10-JUL-1990; GB-015198.
PR
      19-0CT-1990; GB-022845.
PR
      12-NOV-1990; GB-024503.
 PR
      06-MAR-1991; GB-004744.
PR
      15-MAY-1991; GB-010549.
PA
      (CAMB-) CAMBRIDGE ANTIBODY.
PA
      (MEDI-) MED RES COUNCIL.
PΙ
      McCafferty J, Pope AR, Johnson KS, Hoogenboom HRJ, Griffiths AD;
PΙ
      Jackson RH, Holliger KP, Marks JD, Clackson TP, Chiswell DJ;
PI
      Winter GP, Bonnert TP;
DR
      WPI; 92-056862/07.
DR
      P-PSDB; R22582.
PT
      Producing members of specific binding pairs - by expression in
PT
      recombinant host cells with a secreting replicable genetic
 PT
      display package.
PS
      Example 38; Fig 44; 209pp; English.
 CC
      The sequence encodes an antibody scFv fragment directed against 4-
 CC
      hydroxy-3-nitrophenylacetic acid (NP). HV and LV chains of an MAb
CC
      against NP were separately amplified and reassembled to form the
 CC
      construct, which was then ligated into the fd gene III contg. vec-
      tor, fdCAT2, derived from fdTPs/Xh.(See @21095). The clone having
 CC
```

210

```
the scryBl8 sequence (see Q21100) fused in frame to gene III was
 CC
      designated fdCAT2scFvB18. Alternatively the fragment was cloned
 CC
      into fdDOGKan (fdCAT2 with the tet gene replaced by a kanamycin
 CC
      resistance gene) to give fdDOGKanscFvB18, or into the phagemid
 CC
      pHEN1 to create pHEN1-scFvB18. The constructs were used to test
 CC
      the effect of using autator strains to increase the diversity of
 CC
      the cloned genes. The strains NR9046mutD5: NR9046 mutD5::Tn10 and
 CC
      NR9046mutT1: NR9046mutT1::Tn10 were constructed by P1 transduction.
 CC
      After 4 rounds of mutation and screening, 40 phage inserts were
 CC
      sequenced. They each displayed single mutations in 6 different
 CC
      positions, five being in the light chain region. More than 70% of
 CC
      the mutations occurred at positions 724 and 725 changing the first
 CC
      Gly in the J segment (framework 4) to Ser (in 21 cases) or Asp (in
 CC
      3 cases). The mutant shown here (see feature table for details of
 CC
      mutation) occurred three times. The mutant fragments had affinities
 CC
      for NP which were comparable to the wild-type scFv fragment (20nM).
 CC
      N.B. fdCAT2 is also referred to as fd-tet-DOG1 and fdDOG1.
 CC
      See also @21092-100, 103-116, 126-131; @23463, 465-495, 693-719,
 CC
      736-738, and 793-863.
 50
      Sequence
                 770 BP;
                            188 A;
                                      196 C;
                                                214 G;
                                                          172 T;
Initial Score
                      234 Optimized Score =
                                                 279
                                                      Significance = 15.90
Residue Identity =
                                                 287 Mismatches
                      78% Matches
                                                                        66
Gaps
                       11 Conservative Substitutions
                                                                         0
                         X
                                 10
                                           20
                                                     30
                                                               40
                                                                         50
                         GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTC
                         111 | 11111111111111111 | 11111111111 | 11111 | 11111
    TTCTATTCTCACAGTGCACAGGTCCAGCTGCAGCAGTCTGGGGCTTGAGCCTTGTGAAGCCTGGGGCTTCAGTG
            10
                      20 X
                                30
                                          40
                                                    50
                                                              60
           60
                     70
                               80
                                         90
                                                  100
                                                            110
                                                                      120
    AAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAGCAGAGGCCTGAA
    111 111111111 1111111111 11 1 1 1
                                          AAGCTGTCCTGCAAGGCTTCTGGCTACACCTTCACCAGCTACTGGATGCACTGGGTGAAGCAGAGGCCTGGA
          80
                    90
                             100
                                       110
                                                 120
                                                           130
                                                                     140
        130
                  140
                            150
                                      160
                                                170
                                                          180
    CAGGGCCTGGAGTGGATTGGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTC
       11111 111111111111111111111111
                                        11111 1 111111 11 1
    CGAGGCCTTGAGTGGATTGGAAGGATTGATCCTAATAGTGGTGGTACTAAGTACAATGAGAAGTTCAAGAGC
       150
                           170
                 160
                                     180
                                               190
                                                         200
                                                                   210
      200
                210
                          220
                                    230
                                              240
                                                        250
                                                                  260
    AAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGAC
    11111111 | 11 | 1111
                             1111111111111111111111111111111111
    AAGGCCACACTGACTGTAGACAAACCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGACATCTGAGGAC
     220
               230
                         240
                                   250
                                             260
                                                       270
                                                                 280
    270
              280
                         290
                                    300
                                                310
                                                          320
                                                                    330
    ACTGCCGTCTACTACTGTG-CAGA-CGGAATGTGG--GTATCAACGGGATATGCTCTGGACTTCTGGGGCCA
     1111 11111 11 1111 111
                               11 1 11 111 1 11
                                                     11 - 11 - 1 111 111111111
    TCTGCGGTCTATTATTGTGTAAGATACGACTACGGTAGTAGCTAC----TA--CT-TTGACTACTGGGGCCA
   290
             300
                       310
                                 320
                                           330
                                                            340
                                                                      350
      340
                350
                            X
    AGGGACCACGGTCACCGTCTCCTCA
    11111111111111111111111111
    AGGGACCACGGTCACCGTCTCCTCAGGTGGAGGCGGTTCAGGCGGAGGTGGCTCTGGCGGTGGCGGATCCCA
        360
                  370
                            380
                                      390
                                                400
                                                          410
                                                                    420
```

CC

```
ID
     Q15164 standard; DNA; 458 BP.
AC
     015164;
     16-MAR-1992 (first entry)
DT
DE
     VH186 region of anti-nitrophenylacetyl heavy chain Ab gene.
KW
     mouse; murine; antibody; heavy chain; variable region;
K₩
     polymerase chain reaction; ss.
05
     Mus musculus.
PN
     J03247283-A.
PD
     05-NOV-1991.
PF
     29-DEC-1989; 340628.
PR
     29-DEC-1989; JP-340628.
     (MATU ) MATSUSHITA ELEC IND KK.
PA
DR
     WPI; 91-366330/50.
PT
     DNA binding to termini of anti-nitrophenyl:acetyl antibody gene
PT
     - allows specific amplification of variable region in gene by PCR
PS
     Disclosure; Page 2; 3pp; Japanese.
CC
     This sequence corresponds to the region of the heavy chain variable
CC
     region of the murine anti-nitrophenylacetyl IgG antibody which is
CC
     amplified by PCR primers HA and HS.
CC
     See @15159-@15163.
50
     Sequence
             458 BP;
                        112 A;
                                 124 C;
                                          120 G;
                                                  102 T:
Initial Score
                   233 Optimized Score =
                                           277 Significance = 15.83
Residue Identity =
                   78% Matches
                                           283 Mismatches
                                                               72
                     7 Conservative Substitutions
                                                                0
                                                      10
                                                               20
                                               GTCAAACTGCAGCAGTCTGGGG
                                               181 1181181818 88181
   GCTGTATCATGCTCTTCTTGGCAGCAACAGCTACAGGTGTCCACTCCCAGGTCCAACTGCAGCAGCCTGGGG
          20
                   30
                            40
                                    50
                                             60
                                                      70
                                                               80
                                           70
        30
                 40
                          50
                                   60
                                                    80
                                                             90
   CAGAGCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCT
   CTGAGCTTGTGAAGCCTGGGGCTTCAGTGAAGCTGTCCTGCAAGGCTTCTGGCTACACCTTCACCAGCTACT
        90
                100
                         110
                                  120
                                          130
                                                   140
                                                            150
      100
              110
                       120
                                130
                                         140
                                                  150
                                                           160
   ATATGCACTGGGTGAAGCAGAGGCCTGAACAGGGCCTGGAGTGGATTGGAAGGATTGATCCTGCGAGTGGCG
     GGATGCACTGGGTGAAGCAGAGGCCTGGACGAGGCCTTGAGTGGATTGGAAGGATTGATCCTAATAGTGGTG
      160
              170
                       180
                                190
                                         200
                                                  210
                                                           220
    170
             180
                      190
                              200
                                       210
                                                220
                                                         230
   ATACTA AATATGACCCGAAGTTCCAGGTCAAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGC
                230
             240
                     250
                              260
                                       270
                                                280
                                                         290
  240
           250
                    260
                             270
                                     280
                                              290
                                                       300
                                                                310
   TGCAGCTCAGCAGCCTGACATCTGAGGACACTGCCGTCTACTACTGTGCAGACGGAATGTGGGTATCAACGG
   11 1 1 11 1 1111
   TGCAGCTCAGCAGCCTGACATCTGAGGACTCTGCGGTCTATTATTGTGCA---AGA-TACGATTA-CTACGG
  300
           310
                    320
                             330
                                     340
                                                 350
                                                           360
          320
                    330
                             340
                                     350
   GATATGCT-CT-GGACTTCTGGGGCCAAGGGACCACGGTCACCGTCTCCTCA
        TAGTAGCTACTTTGACTACTGGGGCCAAGGCACCACTCTCACAGTCTCCTCAGCCAAAACAACAGCCCCATC
     370
             380
                      390
                               400
                                        410
                                                 420
                                                          430
   GGTCTATCCACTGGCCCCTGT
```

VHI86 region of anti-nitrophenylacetyl heavy chain

GGTCTATCCACTGGCCCCTG

@15164

```
14. US-07-977-702-1 (1-360)
   906227
               VH domain of antibody C against tumour-associated
 ID
     Q06227 standard; DNA; 345 BP.
 AC
     006227;
     22-JAN-1991 (first entry)
 DT
     VH domain of antibody C against tumour-associated antigens.
 DE
 ΚH
     Tumour-associated antigen; murine monoclonal antibody C;
 ΚIJ
     pancreatic carcinoma; metastases; diagnosis; ss.
 08
     Mus musculus.
 PN
     EP-388914-A.
 PD
     26-SEP-1990.
 PF
     21-MAR-1990; 105322.
 PR
     24-MAR-1989; DE-909799.
 PA
     (BEHW ) BEHRINGWERKE AG.
 ΡI
     Bosslet K. Seemann G. Sedlacek HH;
 DR
     WPI; 90-291873/39.
 DR
     P-PSDB; R07320.
PT
     Monoclonal antibodies to tumour associated antigens - used for
PT
     diagnosis of malignant tumours etc.
PS
     Disclosure; Page 13; 18pp; German.
 CC
     Antibody C is produced as described in EP-141079 and binds to
 CC
     pancreatic carcinoma primary tumours and metastases.
CC
     They are useful in tumour diagnosis and therapy.
     See also 906228 for VK of MAb C, 907312-15 for MAb A and B and
 CC
CC
     @06229-30 for MAb D.
SQ
     Sequence 345 BP;
                         87 A;
                                 87 C;
                                          93 G:
                                                  78 T;
                    229 Optimized Score =
Initial Score
                                             245 Significance = 15.53
               =
Residue Identity =
                    71% Matches ·
                                             255 Mismatches
                                        =
                                                                  86
Gaps
                     15 Conservative Substitutions
                                                                   0
             10
                      20
                                30
                                         40
                                                  50
                                                           60
      GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCAGGGGCCTCAGTCAAGTTGTCCTGCACAGCT
      CAGGTCCAACTGCAGCAGTCTGGACCTGAGCTGGTAAAGCCTGGGGCTTCAGTGAAGATGTCCTGCAAGGCT
      X
                    20
          10
                             30
                                      40
                                               50
                                                        60
                                                                  70
  70
                      90
            80
                              100
                                       110
                                                120
                                                         130
   TCTGGCTTCAACATT-AAAGACACCTATATGCACTGGGTGAAGCAGAGGCCTGAACAGGGCCTGGAGTGGAT
   11111 | 1 11111 | 1
                    - 11
                            TCTGGATAC-ACATTCACTACTATGTTATTCACTGGGTGAAACAGAAGCCTGGGCAGGGCCTTGAGTGGAT
         80
                   90
                           100
                                    110
                                              120
                                                       130
                                                                140
          150
                   160
                            170
                                     180
                                               190
                                                        200
                                                                 210
   TGGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCGAAGTTCCAGGTCAAGGCCACTATTACAGC
         TGGATACATTCATCCTTACAATGCTGGTACTGAGTACAATGAGAAGTTCAAAGGCAAGGCCACACTGACTTC
       150
                160
                         170
                                  180
                                            190
                                                     200
                                                              210
        220
                 230
                                   250
                          240
                                             260
                                                      270
   GGACACGTCCTCCAACACACCCTGGCTGCAGCTCAGCAGCCTGACATCTGAGGACACTGCCGTCTACTACTG
    AGACAAATCCTCCAGCACAGCCTACATGGAGCTCAGCAGCCTGACCTCTGAGGACTCTGCGGTCTATTACTG
     220
              230
                       240
                                 250
                                          260
                                                   270
                                                            280
      290
               300
                        310
                                  320
                                           330
                                                    340
                                                             350
   TGCAGACGGAATGTGGGTATCAACGGGATATGCTCTGGACTTCTGGGGCCAAGGGACCACGGTCACCGTCTC
   1 11 1 11 1 1 111 1
                          1 11 11 11 1
                                           111 111 11
   TTCA-ATGGGACGAGGGGT-----GACTA--CTGGGGCC--AAGGGACCACGG--TCACCGTCTCC-TCA
   290
             300
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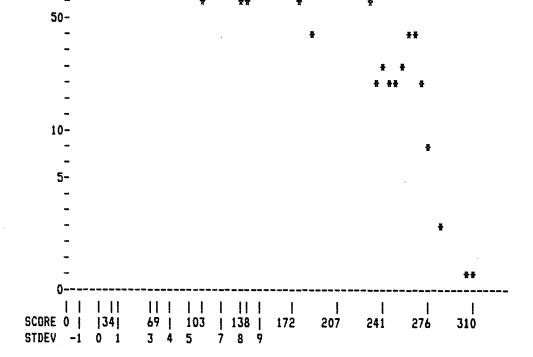
360 CTCA

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15. US-07-977-702-1 (1-360)
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              Genomic to cDNA junction (V47 to VB1-8) DNA
ID
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AC
     N91482:
DT
     28-FEB-1990 (first entry)
DE
     Genomic to cDNA junction (V47 to VB1-8) DNA
К₩
     Human immunodeficiency virus.
05
PN
     ₩08909393-A.
PD
     05-0CT-1989.
PF
     20-MAR-1989; U01152.
PR
     24-MAR-1988; US-173231.
PA
     (IGEN) Igen Inc.
PΙ
     Kenten JH, Casadei J, Well MJ;
DR
     WPI; 89-309634/42.
PT
     New luminescent chimeric proteins - useful in highly sensitive
PT
     immunoassays, eg for HIV
PS
     Disclosure; page 35; 79pp; English.
CC
     This is used in a vector to express the chimeric variable
CC
     4-hydroxy-3-nitrophenyl (NP) antiqen/aequorin protein, in J558L
CC
     myeloma cells or other cells expressing lambda 1 light chain.
CC
     This protein is used as a diagnostic.
50
     Sequence 349 BP; 87 A; 88 C; 96 G; 78 T;
Initial Score
                  228 Optimized Score =
                                         269 Significance = 15.46
                  77% Matches
Residue Identitu =
                                         275 Mismatches
                                                             71
                    7 Conservative Substitutions
Gaps
                                                              0
          10
                  20
                           30
                                    40
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           CAGCAGCCTGGGGCTGAGCTTGTGAAGCCTGGGGCTTCAGTGAAGCTGTCCTGCAAGGCTTCT
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                                               270
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                      AAACCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGACATCTGAGGACTCTGCGGTCTATTATTGTGCA
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                            320
                                      330
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      ---AGA-TACGATTA-CTACGGTAGTAGCTACTTTGACTACTGGGGCCAAGGCACCACTCTCACAGTCTCCT
    280
             290
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                               310
                                        320
                                                330
                                                         340
  360
   CA
   Ш
```

CAG

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O| |O IntelliGenetics
>0 <
FastDB - Fast Pairwise Comparison of Sequences
Release 5.4
Results file us-07-977-702-1.res made by shears on Tue 15 Jun 93 12:27:44-PDT.
Query sequence being compared: US-07-977-702-1 (1-360)
Number of sequences searched:
                                            125798
Number of scores above cutoff:
                                              4369
     Results of the initial comparison of US-07-977-702-1 (1-360) with:
   Data bank : EMBL-NEW 3, all entries
  Data bank : GenBank 75, all entries
  Data bank : GenBank-NEW 3, all entries
  Data bank : UEMBL 34_75, all entries
100000-
U50000-
В
          **
F10000- #
S
E 5000-
9
U
Ε
N
C
E
S 1000-
   500-
   100#
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> 0 <



PARAMETERS

Similarity matrix	Unitary	K-tuple	4
Mismatch penalty	1	Joining penalty	30
Gap penalty	1.00	Window size	32
Gap size penalty	0.33	•	
Cutoff score	0		
Randomization group	0		
Initial scores to save	40	Alignments to save	15
Optimized scores to sav	re 0	Display context	50

SEARCH STATISTICS

Scores:	Mean	Median	Standard Deviation
	28	28	13.45
Times:	CPU 00:30:37.12		Total Elapsed 01:03:42.00

Number of residues: 150464018 Number of sequences searched: 125798 Number of scores above cutoff: 4369

Cut-off raised to 22.
Cut-off raised to 26.
Cut-off raised to 30.
Cut-off raised to 33.
Cut-off raised to 37.
Cut-off raised to 37.
Cut-off raised to 41.
Cut-off raised to 44.
Cut-off raised to 46.
Cut-off raised to 47.
Cut-off raised to 53.
Cut-off raised to 58.
Cut-off raised to 64.

The scores below are sorted by initial score. Significance is calculated based on initial score.

A 100% identical sequence to the query sequence was not found.

				T-:4	0-4		
Seque	nce Name	Description	Length	Init. Score	•	Sig. F	rame
		**** 20 standard deviations	ahovo m		 **		
1.	MMIGGNP2	Mouse mRNA for idiotypic anti		310		20.97	0
	MMCEAHCH	M.musculus gene for anti-CEA		309		20.89	Ö
	MUSIGHNPA	Mouse Ig active H-chain mRNA,		305		20.60	Ö
		**** 18 standard deviations				20.00	•
4.	MUSIGHNP	Mouse Ig active H-chain mRNA,	414	281	314	18.81	0
5.	MMIGGNP1	Mouse mRNA for idiotypic anti	414	281	314	18.81	0
6.	MMV20292B	M.musculus mRNA for VH-gen se	363	276	282	18.44	0
7.	5114902	immunoglobulin heavy chain {n	452	276	281	18.44	0
8.	MMIGVDJAA	M.musculus immunoglobulin hea	452	276	281	18.44	0
9.	MUSIGHNPG	Mouse Ig active H-chain mRNA,	414	273	309	18.22	0
	MMIGGNP8	Mouse mRNA for idiotypic anti	414	273	307	18.22	0
11.	MUSIGHNPB	Mouse Ig active H-chain mRNA,	414	272	307	18.14	0
12.	MMIGGNP3	Mouse mRNA for idiotypic anti	414	272	308	18.14	0
	MMIGG1HCV	M.musculus rearranged mRNA fo	351	271	316	18.07	0
	MUSIGMU4G	Mouse monoclonal antiidiotypi			307	18.07	0
	MUSIGHBB	Mouse Ig active mu-chain VDJ-		271	326	18.07	0
16.	MUSIGHBH	Mouse Ig active H-chain: anti		271	324	18.07	0
17.	MMIGVK1	Mouse mRNA for anti-GAT VH an	501	.271	324	18.07	0
	MUSIGHXO	Mouse Ig germline H-chain gen	970	271	305	18.07	0
	MMIG10VH	Mouse (GAT-specific) subgroup		271	305	18.07	0
	MMIGHVXA	Mouse (hybridoma 3A112) immun			277	18.07	0
21.	MUSIGHVXA	Mouse (hybridoma 3A112) immun		271	277	18.07	0
		**** 17 standard deviations					
	MMV20311	M.musculus mRNA for VH-gen se		270		17.99	0
	MUSIGHADE	Mouse Ig rearranged H-chain g				17.99	0
	MUSIGHEA	Mouse Ig mu-chain active V-re	444	269	322	17.92	0
	MUSIGHDZ	Mouse Ig Mu-chain active V-re	444	269	322	17.92	0
	MUSIGHNPH	Mouse Ig active H-chain mRNA,	417	268	321	17.85	0
	MUSIGHBC	Mouse Ig active H-chain a-NP	511	268	321	17.85	0
	MUSIGHZTA	Mouse Ig geraline H-chain gen				17.77	0
	MUSIGHZP	Mouse Ig germline H-chain gen				17.70	0
	MUSIGHAEM	Mouse Ig heavy-chain mRNA V r				17.70	0
	MUSIGB1H1	Mouse mRNA for immunoglobulin		266		17.70	0
	MMIGB1H1	Mouse mRNA for immunoglobulin		599	271	17.70	0
	MUSIGHZO	Mouse Ig geraline H-chain gen		265		17.62	0
	MUSIGHZN	Mouse Ig geraline H-chain gen		265		17.62	0
	MUSIGHDF	Mouse Ig active mu-chain V-re		265		17.62	0
	MMIGNP4	Mouse mRNA for idiotypic anti		265		17.62	0
	MUSIGHNPE	Mouse Ig active H-chain mRNA,		265		17.62	0
	MMIGGNP6	Mouse mRNA for idiotypic anti		265		17.62	0
	MMIGWM65B	M.musculus WM65 immunoglobuli		265		17.62	0
40.	MMIGHVXZ	Mouse (hybridoma H1-39) immun	1068	265	273	17.62	0

1. US-07-977-702-1 (1-360)

MMIGGNP2 Mouse mRNA for idiotypic anti-NP IgG(1) heavy chai

LOCUS	MMIGGNP2 420 bp RNA ROD 07-MAY-1992
DEFINITION	Mouse mRNA for idiotypic anti-NP IgG(1) heavy chain V-D-J
	(hybridoma 18.1.16)
ACCESSION	X02563 M12744
KEYWORDS	gamma-immunoglobulin; Ig D-segment; Ig heavy chain; immunoglobulin; joining region; signal peptide; variable region.
SOURCE	Mouse
ORGANISM	Mus musculus
	Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;

Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.

REFERENCE 1 (bases 1 to 420)

```
TITLE
            Heavy chain variable region: Multiple gene segments encode
            anti-4-(hydroxy-3-nitrophenyl)acetyl idiotypic antibodies
  JOURNAL
            J. Exp. Med. 161, 1272-1292 (1985)
  STANDARD
            full automatic
 COMMENT
            *source: strain=Balb/c;
            Serological analysis of hybridomaproteins resulting from the immune
            response to the hapten NP reveals NP(a) idiotupes expressed by
            Balb/C mice and NP(b) idiotypes expressed by C57 BL/6 mice. The
            NP(b) family of antibodies consists of 6 subgroups (I-VI) with I-IV
            sharing more determinants than V and VI which appear quite
            distinct.
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                        106 c
                                         97 t
                                110 q
ORIGIN
Initial Score
                    310 Optimized Score =
                                             318 Significance = 20.97
Residue Identity =
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                                             323 Mismatches
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Gaps
                      6 Conservative Substitutions
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                                      50
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                                     60
                                              70
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                       250
                                260
                                         270
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```

Boersch-Supan.M.E., Agarwal.S., White-Scharf.M.E. and

AUTHURS

240

250

260

270

280

290

300

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1 1
                                                                  1 1 111
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                       330
                                340
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   370
             380
                       390
                                 400
                                          410
2. US-07-977-702-1 (1-360)
               M.musculus gene for anti-CEA mAb T84.66 heavy chai
   MMCEAHCH
            MMCEAHCH
LOCUS
                          861 bp
                                   DNA
                                                   ROD
                                                             03-DEC-1992
 DEFINITION M.musculus gene for anti-CEA mAb T84.66 heavy chain V-region
 ACCESSION
            X52769
 KEYWORDS
            CEA; heavy chain; tumor-associated antigen.
 SOURCE
            nouse
   ORGANISM Mus musculus
            Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
            Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
            1 (bases 1 to 861)
            Neumaier, M., Shively, L., Chen, F.S., Gaida, F.J., Ilgen, C.,
   AUTHORS
            Paxton, R.J., Shively, J.E. and Riggs, A.D.
            Cloning of the genes for T86.66, an antibody that has a high
   TITLE
            specificity and affinity for carcinoembryonic antigen, and
            expression of chimeric human/mouse T84.66 genes in myeloma and
            Chinese hamster ovary cells
            Cancer Res. 50, 2128-2134 (1990)
   JOURNAL
   STANDARD full automatic
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                     109..118
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 BASE COUNT
                         208 c
                213 a
                                  203 g
                                          237 t
ORIGIN
Initial Score
                     309 Optimized Score =
                                               316 Significance = 20.89
Residue Identity =
                     88% Matches
                                               322 Mismatches
                                                                      33
Gaps
                      10 Conservative Substitutions
                                                                      0
                                                            10
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                                                        1 111 11 1 111
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             800
                       810
3. US-07-977-702-1 (1-360)
  MUSIGHNPA
              Mouse Ig active H-chain mRNA, V-region (VDJ) from
LOCUS
            MUSIGHNPA
                         421 bp ss-aRNA
                                                          01-SEP-1988
                                                 ROD
 DEFINITION
           Mouse Ig active H-chain mRNA, V-region (VDJ) from anti-NP hybridoma
            18.1.16.
ACCESSION
            M12744
KEYWORDS
            immunoglobulin; immunoglobulin heavy chain; variable region.
SOURCE
            Mouse (BALB/c) hybridoma 18.1.16, cDNA to mRNA.
  ORGANISM
           Mus musculus
            Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
            Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
REFERENCE
            1 (bases 1 to 421)
  AUTHORS
            Boersch-Supan.H.E., Agarwal.S., White-Scharf,M.E. and
            Imanishi-Kari, T.
  TITLE
            Heavy chain variable region Multiple gene segments encode
            anti-4-(hydroxy-3-nitropheny)acetyl idiotypic antibodies
  JOURNAL
            J. Exp. Med. 161, 1272-1292 (1985)
  STANDARD full automatic
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full automatic **FEATURES** Location/Qualifiers

J. Exp. Med. 161, 1272-1292 (1985)

JOURNAL

STANDARD

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                   SSLTSEDTAVYYCASYRYERAWFAYWG@GTLVTVSA"
BASE COUNT
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                       98 c
                               118 q
                                       97 t
ORIGIN
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Initial Score
                   281 Optimized Score =
                                           314 Significance = 18.81
Residue Identity =
                   88% Matches
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                                               Mismatches
                                                               31
Gaps
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5. US-07-977-702-1 (1-360)
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MMIGGNP1 Mouse mRNA for idiotypic anti-NP IgG(1) heavy chai

LOCUS MMIGGNP1 ROD 07-MAY-1992 414 bp RNA DEFINITION Mouse mRNA for idiotypic anti-NP IgG(1) heavy chain V-J (hybridoma 20.1.43)

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SOURCE
  ORGANISM
            Mus ausculus
            Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
            Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
            1 (bases 1 to 414)
 REFERENCE
  AUTHORS
            Boersch-Supan, M.E., Agarwal, S., White-Scharf, M.E. and
            Imanishi-Kari, T.
  TITLE
            Heavy chain variable region: Multiple gene segments encode
            anti-4-(hydroxy-3-nitrophenyl)acetyl idiotypic antibodies
  JOURNAL
            J. Exp. Med. 161, 1272-1292 (1985)
  STANDARD
            full automatic
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            Serological analysis of hybridomaproteins resulting from the immune
            response to the hapten NP reveals NP(a) idiotypes expressed by
            Balb/C mice and NP(b) idiotypes expressed by C57 BL/6 mice. The
            NP(b) family of antibodies consists of 6 subgroups (I-VI) with I-IV
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gamma-immunoglobulin; Ig D-segment; Ig heavy chain; immunoglobulin;

joining region; signal peptide; variable region.

いんりこうごけいい

170

180

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210

220

230

KEYWORDS

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 LOCUS
            MMV20292B
                         363 bp
                                  RNA
                                                 ROD
                                                          23-JUN-1992
DEFINITION M.musculus mRNA for VH-gen sequence of naturally occurring,
            somatically mutated memory B cell
 ACCESSION
            Z12783 X59730
KEYWORDS
            somatic mutation.
 SOURCE
            AOUSE
  ORGANISM Mus musculus
            Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
            Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
            1 (bases 1 to 363)
  AUTHORS
            Schittek,B. and Rajewsky,K.
  TITLE
            Natural occurence and origin of somatically mutated memory B cells
  JOURNAL
            J. Exp. Med. 0, 0-0 (1992)
  STANDARD
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            2 (bases 1 to 363)
  AUTHORS
            Schittek, B.
  TITLE
            Direct Submission
            Submitted (07-JUN-1992) B. Schittek, Institut of Genetics, Weyertal
  JOURNAL
            121, W-5000 Cologne, FRG
  STANDARD full automatic
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            *source: haplotype=IgHb;
            ₹source: stage=Adult;
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X

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                                                        30-NOV-1992
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           cell line 46 transformed with ts mutant of A-MuLV, Genomic, 452 ntl
ACCESSION
           S45712
KEYWORDS
SOURCE
           mice pre-B cell line 46 transformed with ts mutant of A-MuLV
  ORGANISM
           Unclassified.
           Unclassified.
REFERENCE
           1 (bases 1 to 452)
  AUTHORS
           Shirasawa,T., Miyazoe,I., Hagiwara,S., Kimoto,H., Shigemoto,K.,
           Taniguchi, M. and Takemori, T.
  TITLE
           Heavy chain variable (VH) region diversity generated by VH gene
           replacement in the progeny of a single precursor cell transformed
           with a temperature-sensitive mutant of Abelson murine leukemia
           virus.
  JOURNAL
           J. Exp. Med. 176, 1209-1214 (1992)
  STANDARD
           full automatic
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COMMENT
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           This sequence comes from Fig. 2. The authors begin their numbering
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Residue Identity
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Gaps

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     Shirasawa T., Miyazoe I., Hagiwara S., Kimoto H., Shigemoto K.,
RA
     Taniguchi M., Takemori T.;
RT
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RT
     Replacement in the Progeny of a Single Precursor Cell Transformed
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     with a Temperature-sensitive Mutant of Abelson Murine Leukemia
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RP

RA

1-452

Shirasawa T.;

10

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RL
     of Gerontology, 35-2 Sakae-cho, Itabashi-ku, Tokyo, 173, Japan
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πι

310

320

330

340

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9. US-07-977-702-1 (1-360)
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                                                            01-SEP-1988
 DEFINITION
            Mouse Ig active H-chain mRNA, V-region (VDJ) from anti-NP hybridoma
            P3.6.5.
            M12750
 ACCESSION
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            immunoglobulin; immunoglobulin heavy chain; variable region.
 SOURCE
            Mouse (C57BL/6) group V hybridoma P3.6.5, cDNA to mRNA.
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            Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
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            1 (bases 1 to 414)
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            Boersch-Supan,M.E., Agarwal,S., White-Scharf,M.E. and
            Imanishi-Kari, T.
  TITLE
            Heavy chain variable region Multiple gene segments encode
            anti-4-(hydroxy-3-nitropheny)acetyl idiotypic antibodies
            J. Exp. Med. 161, 1272-1292 (1985)
  JOURNAL
  STANDARD full automatic
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10. US-07-977-702-1 (1-360)
   MMIGGNP8
                Mouse mRNA for idiotypic anti-NP IgG(1) heavy chai
LOCUS
            MMIGGNP8
                          414 bp
                                    RNA
                                                   ROD
                                                             07-MAY-1992
 DEFINITION
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            (hubridona P.3.6.5)
 ACCESSION
            X02569 M12750
 KEYWORDS
            gamma-immunoglobulin; Ig D-segment; Ig heavy chain; immunoglobulin;
            joining region; signal peptide; variable region.
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            Mus ausculus
   ORGANISM
            Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
            Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
               (bases 1 to 414)
   AUTHORS
            Boersch-Supan, M.E., Agarwal, S., White-Scharf, M.E. and
            Imanishi-Kari, T.
   TITLE
            Heavy chain variable region: Multiple gene segments encode
            anti-4-(hydroxy-3-nitrophenyl)acetyl idiotypic antibodies
   JOURNAL
            J. Exp. Med. 161, 1272-1292 (1985)
   STANDARD
            full automatic
COMMENT
            *source: strain=C57BL/6;
            Serological analysis of hybridomaproteins resulting from the immune
            response to the hapten NP reveals NP(a) idiotypes expressed by
            Balb/C mice and NP(b) idiotypes expressed by C57 BL/6 mice. The
            NP(b) family of antibodies consists of 6 subgroups (I-VI) with I-IV
            sharing more determinants than V and VI which appear quite
            distinct.
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                     /note="D-region (aa 99-105)"
     misc_feature
                     373..>414
                     /note="J-region (aa 106-119)"
     CDS
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                     /product="idiotypic anti-NP IgG(1) heavy chain V-D-J"
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BASE COUNT
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                         100 c
                                  114 a
                                           98 t
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ORIGIN

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Initial Score
                   273
                        Optimized Score =
                                            307 Significance = 18.22
Residue Identity
                   86% Matches
                                            314 Mismatches
                                                                38
                    10 Conservative Substitutions
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               Mouse Ig active H-chain mRNA, V-region (VDJ) from
LOCUS
           MUSIGHNPB
                        414 bp ss-mRNA
                                               ROD
                                                        01-SEP-1988
DEFINITION
           Mouse Ig active H-chain ARNA, V-region (VDJ) from anti-NP hybridoma
           P5.40.1.
ACCESSION
           M12745
KEYWORDS
           immunoglobulin; immunoglobulin heavy chain; variable region.
SOURCE
           Mouse (C57BL/6) group VI hybridoma P5.40.1, cDNA to mRNA.
  ORGANISM
           Mus musculus
           Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
           Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
REFERENCE
           1 (bases 1 to 414)
  AUTHORS
           Boersch-Supan, M.E., Agarwal, S., White-Scharf, M.E. and
           Imanishi-Kari, T.
  TITLE
           Heavy chain variable region Multiple gene segments encode
           anti-4-(hydroxy-3-nitropheny)acetyl idiotypic antibodies
  JOURNAL
           J. Exp. Med. 161, 1272-1292 (1985)
  STANDARD full automatic
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                               109 g
                                       100 t
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                                            307 Significance = 18.14
Residue Identity =
                   86% Matches
                                            314 Mismatches
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12. US-07-977-702-1 (1-360)
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               Mouse mRNA for idiotypic anti-NP IgG(1) heavy chai
LOCUS
                        414 bp
                                                        07-MAY-1992
DEFINITION Mouse mRNA for idiotypic anti-NP IgG(1) heavy chain V-D-J
           (hybridoma P.5.40.1/group VI)
ACCESSION
           X02564 M12745
KEYWORDS
           gamma-immunoglobulin; Ig D-segment; Ig heavy chain; immunoglobulin;
           joining region; signal peptide; variable region.
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SOURCE

nouse

/note="Ig H-chain V-region"

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REFERENCE
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  AUTHORS
            Boersch-Supan, M.E., Agarwal, S., White-Scharf, M.E. and
            Imanishi-Kari, T.
  TITLE
            Heavy chain variable region: Multiple gene segments encode
            anti-4-(hydroxy-3-nitrophenyl)acetyl idiotypic antibodies
   JOURNAL
            J. Exp. Med. 161, 1272-1292 (1985)
  STANDARD
            full automatic
 COMMENT
            *source: strain=C57 BL/6;
            Serological analysis of hybridomaproteins resulting from the immune
            response to the hapten NP reveals NP(a) idiotypes expressed by
            Balb/C mice and NP(b) idiotypes expressed by C57 BL/6 mice. The
            NP(b) family of antibodies consists of 6 subgroups (I-VI) with I-IV
            sharing more determinants than V and VI which appear quite
            distinct.
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                         101 c
                                 111 q
                                          100 t
 ORIGIN
Initial Score
                     272 Optimized Score =
                                               308 Significance = 18.14
Residue Identity =
                     87% Matches
                                               315 Mismatches
                                                                     37
Gaps
                      10 Conservative Substitutions
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                                                           10
                                                                     20
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Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia; Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.

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13. US-07-977-702-1 (1-360)
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                M.musculus rearranged mRNA for anti-cytochrome c i
LOCUS
             MMIGG1HCV
                          351 bp
                                    RNA
                                                    ROD
                                                              21-NOV-1991
            M.musculus rearranged mRNA for anti-cytochrome c immunoglobulin G1
 DEFINITION
             (IgG1) heavy chain variable region
 ACCESSION
 KEYWORDS
             Anti-cytochrome c immunoglobulin G1; E8 variable heavy chain;
             E8 variable heavy chain D region; E8 variable heavy chain J region;
             E8 variable heavy chain V region; IgG1.
 SOURCE
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  ORGANISM Mus musculus
             Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
             Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
            1 (bases 1 to 351)
   AUTHORS
             Mulvaganam, S.E.
  TITLE
             Direct Submission
   JOURNAL
             Submitted (05-JUL-1991) S.E. Mylvaganam, Dept of Mol Biology, MB4,
             The Scripps Research Inst 10666 N. Torrey Pines Rd, La Jolla, CA
             92307, USA
   STANDARD
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 REFERENCE
            2 (bases 1 to 351)
   AUTHORS
             Mylvaganam,S.E., Paterson,Y., Kaiser,K., Bowdish,K. and
             Getzoff,E.D.
   TITLE
             Biochemical Implications from the Variable Gene Sequences of an
             Anti-cytochrome c Antibody and Crystallographic Characterization of
             its Antigen-binding Fragment in Free and Antigen-complexed Forms
   JOURNAL
             J. Mol. Biol. 221, 455-462 (1991)
   STANDARD full automatic
 COMMENT
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             *source: cell_type=myeloma;
             *source: cell_line=CA4-1;
             E8 is a monoclonal antibody of the anti-cytochrome c immunoglobulin
             G1 (IgG1) that binds horse cytochrome c. E8 variable light chain -
             X60684
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                                           316 Significance = 18.07
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                                           323 Mismatches
Gaps
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14. U5-07-977-702-1 (1-360)
   MUSIGMU4G
               Mouse monoclonal antiidiotypic antibody IgM VDJ-re
LOCUS
           MUSIGMU4G
                        354 bp ss-mRNA
                                               ROD
                                                        16-JUL-1992
DEFINITION Mouse monoclonal antiidiotypic antibody IgM VDJ-region mRNA.
ACCESSION
           M83722
KEYWORDS
           diversity region; immunoglobulin heavy chain;
           immunoglobulin mu-chain; joining region;
           monoclonal antiidiotypic antibody; variable region.
SOURCE
           Mus musculus hubridoma cDNA to mRNA.
  ORGANISM Mus musculus
           Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
           Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
REFERENCE
           1 (bases 1 to 354)
  AUTHORS
           Taub, R., Hsu, J.-C., Garsky, V.M., Hill, B.L., Erlanger, B.F. and
           Kohn, L.D.
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anti-idiotypic antibodies against the thyrotropin (TSH) receptor
            are similar to TSH and inhibit TSH-incresed cAMP production in
           FRTL-5 thyroid cells
   JOURNAL
           J. Biol. Chem. 267, 5977-5984 (1992)
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Residue Identity =
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15. US-07-977-702-1 (1-360)
   MUSIGHBB
               Mouse Ig active mu-chain VDJ-region mRNA from hybr
 LOCUS
           MUSIGHBB
                        390 bp ss-ARNA
                                               ROD
                                                        30-JUN-1987
 DEFINITION Mouse Ig active mu-chain VDJ-region mRNA from hybridoma 87.92.6.
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constant region; diversity region; immunoglobulin heavy chain;

ACCESSION

KEYWORDS

M13832

reptide sequences from the hypervariable regions of two monoclonal

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variable region.
 SOURCE
           Mouse (Balb/c) hybridoma 87.92.6 (mu,kappa), cDNA to mRNA.
  ORGANISM Mus musculus
           Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
           Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
           1 (bases 1 to 390)
  AUTHORS
           Bruck, C., Co, M.S., Slaoui, M., Gaulton, G.N., Smith, T., Fields, B.N.,
           Mullins, J.I. and Greene, M.I.
  TITLE
           Nucleic acid sequence of an internal image-bearing monoclonal
           anti-idiotype and its comparison to the sequence of the external
           antigen
  JOURNAL
           Proc. Natl. Acad. Sci. U.S.A. 83, 6578-6582 (1986)
  STANDARD full automatic
                   Location/Qualifiers
FEATURES
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                    <1..30
                   /codon_start=1
                   /note="Immunoglobulin mu-chain signal peptide"
                   31..>390
     mat_peptide
                   /codon_start=1
                   /note="Immunoglobulin mu-chain"
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                   327..328
                   /note="V-region end/D-region start"
     misc_recomb
                   342..343
                   /note="D-region end/J-region start"
     CDS
                   <1..>390
                   /note="Immunoglobulin mu-chain precursor"
                   /codon start=1
                   translation="LMAVVTGVNSEVQLQQSGAELVKPGASVKLSCTASGFNIKDTYM/
                   HWVK@RPE@GLEWIGRIDPANGNTKYDPKF@GKATITADTSSNTAYL@LSSLTSEDTA
                   VYYCARGGLRRGYAMDYWG@GTSVTVSS"
BASE COUNT
               100 a
                        96 c
                               111 g
ORIGIN
           44 bp upstream of PstI site; chromosome 12.
Initial Score
                   271 Optimized Score =
                                           326 Significance = 18.07
Residue Identity =
                   91% Matches
                                           330 Mismatches
                                                                25
                     7 Conservative Substitutions
Gaps
                                                                 0
                                        10
                                                 20
                                                          30
                                 GTCAAACTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCA
                                    CTGATGGCAGTGGTTACAGGGGTCAATTCAGAGGTTCAGCTGCAGCAGTCTGGGGCAGAGCTTGTGAAGCCA
          10
                   20
                            30
                                X
                                     40
                                              50
                                                       60
                                                                70
  40
           50
                    60
                             70
                                      80
                                               90
                                                       100
                                                                110
   GGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAG
   GGGGCCTCAGTCAAGTTGTCCTGCACAGCTTCTGGCTTCAACATTAAAGACACCTATATGCACTGGGTGAAG
        80
                 90
                          100
                                  110
                                           120
                                                    130
                                                              140
         120
                 130
                          140
                                   150
                                            160
                                                     170
                                                              180
   CAGAGGCCTGAACAGGGCCTGGAGTGGATTGGAAGGATTGATCCTGCGAGTGGCGATACTAAATATGACCCG
   CAGAGGCCTGAACAGGGCCTGGAGTGGATTGGAAGGATTGATCCTGCGAATGGTAATACTAAATATGACCCG
      150
               160
                        170
                                 180
                                          190
                                                   200
                                                            210
       190
               200
                        210
                                  220
                                          230
                                                    240
   AAGTTCCAGGTCAAGGCCACTATTACAGCGGACACGTCCTCCAACACAGCCTGGCTGCAGCTCAGCAGCCTG
   AAGTTCCAGGGCAAGGCCACTATAACAGCAGACACATCCTCCAACACAGCCTACCTGCAGCTCAGCAGCCTG
    220
             230
                      240
                               250
                                        260
                                                 270
                                                          280
     260
              270
                       280
                                 290
                                          300
                                                    310
                                                             320
   ACATCTGAGGACACTGCCGTCTACTACTGTGC-AGACGGAATGTGGGTATCAAC-GGGATATGCTCTGGACT
```

ACATCTGAGGACACTGCCGTCTATTACTGTGCTAGAGGG-----GGATTACGACGGGGGTATGCTATGGACT

immunuytooutin mu-chain; joining region; processe**d gene**;

```
TCTGGGGCCAAGGGACCACGGTCACCGTCTCCTCA
     111111 11111 111 1 111111111111111
    ACTGGGGTCAAGGAACCTCAGTCACCGTCTCCTCA
      360
                370
                          380
 Seq. 2
> 0 <
0| | 0 IntelliGenetics
>0 <
FastDB - Fast Pairwise Comparison of Sequences
Release 5.4
Results file us-07-977-702-2-ngs.res made by shears on Tue 15 Jun 93 11:11:14-PDT.
Query sequence being compared:US-07-977-702-2 (1-318)
Number of sequences searched:
                                             25646
Number of scores above cutoff:
                                              4637
      Results of the initial comparison of US-07-977-702-2 (1-318) with:
   Data bank : N-GeneSeq 10, all entries
 10000-#
U 5000-
F 1000-
  500*
  100-
    50-
                         香香
    10-
```

330

330

Ħ В Ε R

0

S

9 U Ε N C Ε 340

350

340

PARAMETERS

Similarity matrix	Unitary	K-tuple	4
Mismatch penalty	1	Joining penalty	30
Gap penalty	1.00	₩indow size	32
Gap size penalty	0.33		
Cutoff score	0		
Randomization group	0		
Initial scores to save	40	Alignments to save	15
Optimized scores to sav	e 0	Display context	50

SEARCH STATISTICS

Scores:	Mean	Median	Standard Deviation		
	17	14	12.85		
Times:	CPU 00:03:01.00		Total Elapsed 00:06:12.00		

Number of residues: 14371384 Number of sequences searched: 25646 Number of scores above cutoff: 4637

Cut-off raised to 8. Cut-off raised to 15. Cut-off raised to 22. Cut-off raised to 26. Cut-off raised to 29.

The scores below are sorted by initial score. Significance is calculated based on initial score.

A 100% identical sequence to the query sequence was not found.

The list of best scores is:

Sequence Name	Description	Length	Init. Score	•	Sig.	Frame
	**** 19 standard deviations	above me	an **	 ++		
1. 004265	Encodes Colon Cancer monoclon	381	263	265	19.14	0
	**** 18 standard deviations	above me	an **	F#		
2. 028254	VL FRP5.	407	260	263	18.91	0
3. 028653	DNA encoding 1E6 kappa light	327	257	260	18.68	0
4. 928262	Fv(FRP5)-ETA fusion gene.	2011	257	260	18.68	0
5. 028257	Fv(FRP5)-phoA recombinant ant	2233	257	260	18.68	0
6. 004592	Sequence encoding region of h	324	256	257	18.60	0
7. N90813	DNA encoding light chain vari	324	256	257	18.60	0
8. 0 27132	Light chain variable region c	387	256	257	18.60	0
9. 028256	Fv heavy chain/light chain va	748	256	259	18.60	0
10. @05557	Sequence encoding variable re	411	255	259	18.52	0
11. N40022	cDNA insert of pK17G4 encodin	882	252	254	18.29	0
12. 004264	Encodes Colon Cancer monoclon	685	250	253	18.13	0

```
mmma encooling kappa anti-carc
                    **** 16 standard deviations above mean ****
  14. @25592
                   Encodes 4D5 Fab IgG antibody
                                                    2178
                                                           234
                                                                 237 16.89
  15. 028747
                   Anti-CEA specific light chain
                                                     387
                                                           227
                                                                 257
                                                                      16.34
                    **** 15 standard deviations above mean ****
  16. N81636
                   Anti-pseudomonas aeruginosa h
                                                     324
                                                           216
                                                                 219
                                                                      15.49
  17. N81638
                   Anti-pseudomonas aeruginosa h
                                                     671
                                                           216
                                                                 220
                                                                      15.49
  18. 010946
                   Feline immunoglobulin kappa c
                                                     618
                                                          214
                                                                 221
                                                                      15.33
  19. @11191
                   V region gene JP2gL411 from a
                                                     618
                                                           214
                                                                 221 15.33
  20. @04654
                   Plasmid pBT111 encoding antib
                                                    5238
                                                           213
                                                                 214 15.25
  21. 029150
                   WN1 222-5 antibody Light chai
                                                     322
                                                           210
                                                                 215 15.02
                    **** 14 standard deviations above mean ****
  22. 012017
                   Sequence encoding mouse MAb 1
                                                     402
                                                           208
                                                                 216 14.86
  23. @12061
                   Sequence encoding light (kapp
                                                     402
                                                           208
                                                                 216
                                                                     14.86
  24. 029151
                   WN1 58-9 antibody Light chain
                                                     321
                                                           204
                                                                 213 14.55
  25. 015114
                   IL-2 chimeric antibody light
                                                     432
                                                           204
                                                                 211 14.55
  26. 023566
                   Rat CD4 antibody light chain
                                                     620
                                                           204
                                                                 209 14.55
  27. 005556
                   Sequence encoding variable re
                                                           200
                                                     419
                                                                 210 14.24
  28. N90672
                   DNA sequence encoding uproces
                                                     420
                                                           200
                                                                 208 14.24
  29. 015113
                   IL-2 chimeric antibody light
                                                     432
                                                           200
                                                                 207
                                                                      14.24
  30. 008608
                                                                 208
                   ME4 Light Chain V Region (mou
                                                     444
                                                           200
                                                                      14.24
  31. 012633
                   CD4-specific CDR-grafted ligh
                                                     754
                                                           200
                                                                 208 14.24
  32. N90500
                   Light chain antibody variable
                                                     806
                                                           198
                                                                 206
                                                                      14.09
  33. 023567
                   Reshaped CAMPATH-1 antibody l
                                                     748
                                                           198
                                                                 206 14.09
  34. @20381
                   Sequence encoding the shorten
                                                     819
                                                           198
                                                                 208
                                                                     14.09
  35. 023568
                   Reshaped CD4 antibody light c
                                                     748
                                                           197
                                                                 205
                                                                     14.01
  36. N90300
                   Insert coding for light chain
                                                    1395
                                                           197
                                                                 207
                                                                      14.01
                    *** 13 standard deviations above mean ***
  37, 011848
                                                          194
                   Human anti-placental alkaline
                                                     324
                                                                 204
                                                                      13.77
                   FabD1.3 in pUC19.
  38. 021097
                                                    1526
                                                           194
                                                                 203
                                                                      13.77
  39. 011846
                   Anti-placental alkaline phosp
                                                     324
                                                           193
                                                                 203 13.70
                                                                              0
  40. N30165
                   Sequence encoding the leader,
                                                     450
                                                           192
                                                                 199 13.62
1. US-07-977-702-2 (1-318)
   004265
                Encodes Colon Cancer monoclonal antibody CC92 ligh
 ID
      Q04265 standard; DNA; 381 BP.
 AC
      004265;
 DT
      13-SEP-1990 (first entry)
      Encodes Colon Cancer monoclonal antibody CC92 light chain variable region
 DE
 KH
      chimeric antibodies; TAG72; light chain variable region;
 KW
      heavy chain variable region; ss
      synthetic.
 OS
 PN
      EP-365997-A.
 PD
      02-MAY-1990.
 PF
      18-OCT-1989; 119361.
 PR
      19-OCT-1988; US-259943.
 PA
      (DOWC) Dow Chemical Co.
 PΙ
      Mezes P, Gourlie B, Rixon M;
 DR
      WPI; 90-133521/18.
 DR
      P-PSDB; R04388.
 PT
      Chimeric antibodies against TAG72 and conjugate to provide imaging
 PT
      markers and therapeutic tools
```

Initial Score = 263 Optimized Score = 265 Significance = 19.14
Residue Identity = 83% Matches = 266 Mismatches = 52
Gaps = 1 Conservative Substitutions = 0

90 C;

The polypeptide encoded by this sequence forms part of a chimera. The

other components are a heavy chain variable region and human-derived

when administered to human patients because they are of human origin.

97 G;

constant light and heavy chain regions. The variable regions have high affinity for TAG72. The constant regions reduce the side-effects

PS

CC

CC

CC

CC

CC

SQ

Disclosure; pp; English.

See also 004258-64 and 004266-9.

90 A;

381 BP;

```
AG-TATTGTGATGACCCAGACT
                                                                                                         AGATCCAGGTCCTTTGTATTCGTGTTTCTCTGGTTGTCTGGTGTTGACGGAGACATTGTGATGACCCAGTCT
         10
                                              30
                                                                40
                                                                                  50
                                                                                                     60
                                                                                                                       70
                                      40
                                                         50
                                                                           60
                                                                                             70
                                                                                                                80
                                                                                                                                  90
       CCCAAATTCCTGCTTGTTTCAGCAGGAGACAGGGTTACCATAACCTGCAAGGCCAGTCAGAGTGTGACTAAT
                                        1111 | 11111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 1111 | 11
       CACAAATTCATGTCCACATCAGTAGGAGACAGGGTCAGCATCACCTGCAAGGCCAGTCAGGATGTGAGTAGT
                       90
                                        100
                                                           110
                                                                             120
                                                                                               130
                                                                                                                  140
              100
                                 110
                                                   120
                                                                     130
                                                                                        140
                                                                                                           150
                                                                                                                             160
       GATGTAGCTTGGTACCAACAGAAGCCAGGGCAGTCTCCTAAACTGCTGATATATTATGCATCCAATCGCTAC
       GCTGTAGGGTGGTTTCAACAGAAACCAGGACAATCTCCTAAATTACTGATTTATTCGGCATCCTACCGGTAT
                  160
                                    170
                                                       180
                                                                         190
                                                                                           200
                                                                                                              210
                                                                                                                                220
           170
                             180
                                               190
                                                                  200
                                                                                    210
                                                                                                       220
                                                                                                                         230
       ACTGGAGTCCCTGATCGCTTCACTGGCAGTGGATATGGGACGGATTTCACTTTCACCATCAGCACTGTGCAG
       230
                                 240
                                                   250
                                                                     260
                                                                                        270
                                                                                                          280
                                                                                                                             290
       240
                         250
                                            260
                                                              270
                                                                                 280
                                                                                                   290
                                                                                                                      300
       GCTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTACACGTTCGGAGGGGGGACCAAG
       1111 11111111 1 11111111
       GCTGAAGACCTGGCAGTTTATTACTGTCAGCAACATTATAGTAGTCCGCTCACGTTCGGTGCTGGGACCAAG
           300
                             310
                                               320
                                                                  330
                                                                                    340
                                                                                                       350
                                                                                                                         360
   310
                     X
       CTGGAGATC
       111111
       CTGGAGCTGAAACT
       370
                     X 380
2. US-07-977-702-2 (1-318)
     028254
                             VL FRP5.
 ID
           928254 standard; DNA; 407 BP.
 AC
           028254;
 DT
          11-FEB-1993 (first entry)
 DE
           VL FRP5.
 KW
           Minigene; pMZ18/1; monoclonal antibody; kappa; light chain;
 K₩
           variable region; pWW31; ss.
 05
           Mus musculus.
 PN
           EP-502812-A.
 PD
           09-SEP-1992.
 PF
           27-JAN-1992; 810056.
 PR
          05-FEB-1991; EP-810079.
 PA
           (CIBA ) CIBA GEIGY AG.
 PI
           Groner B, Hardman N, Harwerth I, Hunes NE, Wels WS;
 PI
           Zwickl M;
 DR
           WPI; 92-302096/37.
 PT
           Recombinant antibodies directed to growth factor receptor C-erbB-2 -
 PT
           for diagnosing and treating tumours expressing C-erbB-2 e.g. breast
 PT
           or ovarian tumours
 PS
           Disclosure; Page 29; 67pp; English.
 CC
           The sequence given a portion of the plasmid pMZ18/1 and encodes the
 CC
           kappa light chain variable (V) domain of the monoclonal antibody FRP5.
 CC
           The V-region was excised from this plasmid and cloned into plasmid
 CC
           pWW31 which was used in the construction of a recombinant antibody
```

which is directed to the extracellular domain of the growth factor

CC

CC

c-erbB-2.

X

10

```
Initial Score
                   260 Optimized Score =
                                           263 Significance = 18.91
Residue Identity =
                   82% Matches
                                           264 Mismatches
                     1 Conservative Substitutions
Gaps
                                                                 0
                           X
                                   10
                                            20
                                                     30
                                                              40
                           AG-TATTGTGATGACCCAGACTCCCAAATTCCTGCTTGTTTCAGC
                            TCTAGTCACTGGATGGTGGGAAGATGGAGACATTGTGATGACCCAGTCTCACAAATTCCTGTCCACTTCAGT
                            30
                                     40
                                              50
                                                       60
                                                                70
       50
                60
                         70
                                  80
                                           90
                                                   100
                                                            110
   AGGAGACAGGGTTACCATAACCTGCAAGGCCAGTCAGAGTGTGACTAATGATGTAGCTTGGTACCAACAGAA
   AGGAGACAGGGTCAGCATCACCTGCAAGGCCAGTCAGGATGTGTATAATGCTGTTGCCTGGTATCAACAGAA
         80
                  90
                          100
                                  110
                                           120
                                                    130
                                                             140
    120
             130
                      140
                               150
                                        160
                                                 170
                                                          180
   GCCAGGGCAGTCTCCTAAACTGCTGATATATTATGCATCCAATCGCTACACTGGAGTCCCTGATCGCTTCAC
    ACCAGGACAATCTCCTAAACTTCTGATTTACTCGGCATCCTCCCGGTACACTGGAGTCCCTTCTCGCTTCAC
      150
               160
                        170
                                 180
                                          190
                                                   200
                                                           210
  190
           200
                    210
                             220
                                      230
                                               240
                                                        250
                                                                 260
   TGGCAGTGGATATGGGACGGATTTCACTTTCACCATCAGCACTGTGCAGGCTGAAGACCTGGCAGTTTATTT
   TGGCAGTGGCTCTGGGCCGGATTTCACTTTCACCATCAGCAGTGTGCAGGCTGAAGACCTGGCAGTTTATTT
    220
                               250
             230
                      240
                                        260
                                                 270
                                                          280
         270
                  280
                           290
                                    300
                                             310
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   CTGTCAGCAACATTTTCGTACTCCATTCACGTTCGGCTCGGGGACAAAATTGGAAATAAAACGGGCTGATGC
  290
           300
                    310
                             320
                                      330
                                               340
                                                      X 350
                                                                 360
   TGCACCAACTGTATCCATCTTCCCACCATCCAGTGA
         370
                  380
                           390
3. US-07-977-702-2 (1-318)
  028653
              DNA encoding 1E6 kappa light chain variable region
ID
     Q28653 standard; DNA; 327 BP.
AC
     028653;
DT
     17-FEB-1993 (first entry)
DE
     DNA encoding 1E6 kappa light chain variable region.
K₩
     Chimeric; heavy chain; variable region; monoclonal antibody;
K₩
     lymphocyte function associated antigen-3; LFA-3; inflammation;
KW
     autoimmune disease; immunomodulation; systemic lupus erythematosus;
KW
     rheumatoid arthritis; thuroiditis; ss.
05
     Homo sapiens.
PN
     EP-503646-A.
PD
     16-SEP-1992.
PF
     12-MAR-1992; 104318.
PR
     12-MAR-1991; US-667975.
PA
     (BIOJ ) BIOGEN INC.
PΙ
     Chisholm PL, Sato VL, Wallner BP;
     WPI; 92-309758/38.
 DR
 DR
     P-PSDB; R27144.
PT
     Monoclonal antibodies active against lymphocyte function
PT
     associated antigen-3 - for treating inflammation and auto:immune
PT
     diseases, and for detecting LFA-3 protein in a sample
PS
     Claim 16; Page 14; 30pp; English.
 CC
     DNA sequences obtd. from the hybridoma cell line 1E6-2C12 (ATCC HB
```

10693) encoding the light and heavy chain regions of 1E6 were

CC

10/ Ci

```
CC
     the light chain variable region were MKVP7 and 360-41, and the heavy
 CC
     chain primers were VHO1 and VHO2. The PCR prods. were
 CC
     electrophoresed on agarose gel and the DNA excised and inserted into
 CC
     plasmids. The light and heavy chain regions may be used to
 CC
     construct a monoclonal antibody active against LFA-3 which blocks
 CC
     adhesion of LFA-3 expressing cells to lymphocytes. The antibody may
 CC
     be used in the treatment and diagnosis of acute and chronic
 CC
     inflammation, autoimmune diseases and for immunomodulation including
 CC
     systemic lupus erythematosus, rheumatoid arthritis and thyroiditis.
 CC
     See also 028654-8.
 SQ
     Sequence
               327 BP;
                          89 A;
                                  82 C;
                                           83 G;
                                                   73 T;
Initial Score
                    257 Optimized Score =
                                             260 Significance = 18.68
Residue Identity =
                    81% Matches
                                              260 Mismatches
                                                                   58
Gaps
                      O Conservative Substitutions
                                                                    0
           10
                    20
                              30
                                       40
                                                50
                                                          60
                                                                   70
   AGTATTGTGATGACCCAGACTCCCAAATTCCTGCTTGTTTCAGCAGGAGACAGGGTTACCATAACCTGCAAG
      11111 1111111 1111111 1 11
                                      1 1111 11111 11111 111 1 111111111
   AACATTGTAATGACCCAATCTCCCAAATCCATGTCCATGTCAGTAGGAGAGGGGTCACCTTGACCTGCAAG
           10
                    20
                              30
                                       40
                                                50
                                                                   70
                                                          60
         80
                  90
                           100
                                    110
                                              120
                                                       130
                                                                140
   GCCAGTCAGAGTGTGACTAATGATGTAGCTTGGTACCAACAGAAGCCAGGGCAGTCTCCTAAACTGCTGATA
    GCCAGTGAGAATGTGGTTACTTATGTTTCCTGGTATCAACAGAAACCAGAGCAGTCTCCTAAACTGCTCATA
         80
                  90
                           100
                                    110
                                             120
                                                       130
                                                                140
      150
                160
                         170
                                  180
                                            190
                                                     200
                                                              210
   TATTATGCATCCAATCGCTACACTGGAGTCCCTGATCGCTTCACTGGCAGTGGATATGGGACGGATTTCACT
         TACGGGGCATCCAACCGGTACACTGGGGTCCCCGATCGCTTCACAGGCAGTGGATCTGCAACAGATTTCACT
      150
                160
                         170
                                  180
                                            190
                                                     200
                                                              210
    220
              230
                       240
                                250
                                          260
                                                   270
                                                            280
   TTCACCATCAGCACTGTGCAGGCTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTAC
    1111 | 111 | 111 | 111111
   CTGACCATCAGCAGTGTGCAGGCTGAAGACCTTGCAGATTATCACTGTGGACAGGGTTACAGCTATCCGTAC
    220
              230
                       240
                                250
                                          260
                                                   270
                                                            280
  290
            300
                     310
                              X
   ACGTTCGGAGGGGGGACCAAGCTGGAGATC
   ACGTTCGGAGGGGGGACCAAGCTGGAAATAAAACGGGCT
  290
            300
                     310
                              320
4. US-07-977-702-2 (1-318)
  928262
              Fv(FRP5)-ETA fusion gene.
 ID
     028262 standard; DNA; 2011 BP.
 AC
     928262;
 DT
     11-FEB-1993 (first entry)
 DE
     Fv(FRP5)-ETA fusion gene.
 Κ₩
     Monoclonal antibody; light chain; heavy chain; tumour; c-erbB-2;
 K₩
     variable region; ETA; ss.
 05
     Pseudomonas aeruginosa PAK.
FH
     Keu
                    Location/Qualifiers
FT
     sig_peptide
                    1..63
FT
     /*tag= a
 FT
     /label= ompA_signal_peptide
FT
     misc_RNA
                    64..87
 FT
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 FT
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amplified for cloning and sequencing using PCK. The PCK primers for

しし

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misc RNA

97..453

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 PN
     EP-502812-A.
 PD
     09-SEP-1992.
 PF
     27-JAN-1992; 810056.
 PR
     05-FEB-1991; EP-810079.
PA
     (CIBA ) CIBA GEIGY AG.
 PΙ
     Groner B, Hardman N, Harwerth I, Hynes NE, Wels WS;
PΙ
     Zwickl M;
 DR
     WPI; 92-302096/37.
 DR
     P-PSDB; R26982.
PT
     Recombinant antibodies directed to growth factor receptor C-erbB-2 -
 PT
     for diagnosing and treating tumours expressing C-erbB-2 e.g. breast
 PT
     or ovarian tumours
PS
     Disclosure; Page 47-52; 67pp; English.
 CC
     The sequences given in 028261-3 encode part of the exotoxin A (ETA)
 CC
     sequence corresponding to positions 1574-1747 of the full exotoxin A
 CC
     sequence. These sequences were used in the construction of
 CC
     Fv(FRP5)-ETA fusion genes. The ETA sequence was used as a marker gene
 CC
     so that E. coli transformed with the fusion gene could be identified.
     The fusion genes were expressed in E. coli and the antibody was
 CC
 CC
     extracted. This recombinant antibody can be used for the qualitative
     and quantitative determination of c-erbB-2. This can be used for
 CC
CC
     monitoring or in-vivo localisation of tumours overexpressing c-erbB-2.
50
               2011 BP;
     Sequence
                           383 A;
                                    653 C;
                                              620 G;
Initial Score
                    257 Optimized Score =
                                              260 Significance = 18.68
Residue Identity =
                    81% Matches
                                              260 Mismatches
                                                                   58
Gaps
                      O Conservative Substitutions
                                                                    0
                                                          10
                                                                   20
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                                                          1 1111111 111
   CCTCTGGCGGTGGCGGTTCTGGTGGCGGTGGCCGGTGGCGGTTCTGACATCCAGCTGACCCAGTCTC
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            460
                     470
                               480
                                        490
                                                  500
                                                           510
                                                                    520
         30
                  40
                            50
                                     60
                                               70
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                  ACAAATTCCTGTCCACTTCAGTAGGAGACAGGGTCAGCATCACCTGCAAGGCCAGTCAGGATGTGTATAATG
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                                      560
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                                                         580
                                                                  590
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                110
                         120
                                  130
                                            140
                                                     150
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                 610
                           620
                                    630
                                              640
                                                       650
                                                                 660
    170
              180
                       190
                                200
                                          210
                                                   220
                                                             230
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    CTGGAGTCCCTTCTCGCTTCACTGGCAGTGGCTCTGGGCCGGATTTCACTTTCACCATCAGCAGTGTGCAGG
      670
               680
                         690
                                  700
                                            710
                                                     720
                                                               730
```

ГΙ

```
CJV
                      cov
                                C/V
                                          ZÖV
    CTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTACACGTTCGGAGGGGGGGACCAAGC
    CTGAAGACCTGGCAGTTTATTTCTGTCAGCAACATTTTCGTACTCCATTCACGTTCGGCTCGGGGACAAAAT
    740
              750
                        760
                                  770
                                                      790
                                                               800
          X
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    TGGAGATCAAAGCTCTAGAGGGCGGCAGCCTGGCCGCGCTGACCGCGCACCAGGCCTG
          X 820
   810
                      830
                                          850
5. US-07-977-702-2 (1-318)
   928257
               Fv(FRP5)-phoA recombinant antibody.
 ID
     Q28257 standard; DNA; 2233 BP.
     928257;
 AC
 DT
     11-FEB-1993 (first entry)
     Fv(FRP5)-phoA recombinant antibody.
 DE
     Monoclonal antibody; light chain; heavy chain; tumour; phoA;
 KW
 KW
     variable region; alkaline phosphatase; c-erbB-2; ss
 05
     Synthetic.
 FH
     Keu
                     Location/Qualifiers
FT
     CDS
                     23..2158
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 FT
FT
     sig_peptide
                     23..85
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FT
FT
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     Misc_RNA
FT
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     /label= phoA_coding_region
PN
     EP-502812-A.
PD
     09-SEP-1992.
PF
     27-JAN-1992; 810056.
PR
     05-FEB-1991; EP-810079.
PA
     (CIBA ) CIBA GEIGY AG.
     Groner B, Hardman N, Harwerth I, Hynes NE, Wels WS;
PΙ
PΙ
     Zwickl M;
     WPI; 92-302096/37.
DR
DR
     P-PSDB; R26980.
PT
     Recombinant antibodies directed to growth factor receptor C-erbB-2 -
PT
     for diagnosing and treating tumours expressing C-erbB-2 e.g. breast
PT
     or ovarian tumours
     Disclosure; Page 34-40; 67pp; English.
PS
CC
     The sequence given encodes the single chain recombinant antibody
CC
     desiganted Fv(FRP5)-phoA. The alkaline phosphatase gene (phoA) was
CC
     used as a marker gene so that E. coli transformed with the fusion gene
CC
     could be identified. The fusion gene was expressed in E. coli and the
CC
     antibody was extracted. This recombinant antibody can be used for the
CC
     qualitative and quantitative determination of c-erbB-2. This can be
CC
     used for monitoring or in-vivo localisation of tumours overexpressing
CC
     c-erbB-2.
SQ
     Sequence
               2233 BP;
                            566 A;
                                      573 C;
                                               606 G;
                                                         488 T;
Initial Score
                     257 Optimized Score =
                                               260 Significance = 18.68
Residue Identity =
                     81% Natches
                                               260 Mismatches
```

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10
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                                               AGTATTGTGATGACCCAGACTC
                                                      1 11111111 111
   450
                  460
                           470
                                    480
                                             490
                                                      500
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                 40
                          50
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                                           560
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               110
                        120
                                130
                                         140
                                                  150
                                                           160
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    CTGTTGCCTGGTATCAACAGAAACCAGGACAATCTCCTAAACTTCTGATTTACTCGGCATCCTCCCGGTACA
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               600
                        610
                                620
                                         630
                                                  640
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             180
                      190
                               200
                                        210
                                                220
                                                         230
   CTGGAGTCCCTGATCGCTTCACTGGCAGTGGATATGGGACGGATTTCACTTTCACCATCAGCACTGTGCAGG
   CTGGAGTCCCTTCTCGCTTCACTGGCAGTGGCTCTGGGCCGGATTTCACTTTCACCATCAGCAGTGTGCAGG
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             670
                      680
                               690
                                        700
                                                710
                                                         720
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           250
                    260
                             270
                                      280
                                               290
                                                       300
                                                                310
   CTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTACACGTTCGGAGGGGGGACCAAGC
   CTGAAGACCTGGCAGTTTATTTCTGTCAGCAACATTTTCGTACTCCATTCACGTTCGGCTCGGGGACAAAAT
  730
           740
                    750
                             760
                                      770
                                               780
                                                       790
                                                                800
         X
   TGGAGATC
   11111111
   TGGAGATCAAAGCTCTAGAGCCTGTTCTGGAAAACCGGGCTGCTCAGGGCGATATTAC
         810
                  820
                           830
                                    840
                                             850
6. US-07-977-702-2 (1-318)
  004592
              Sequence encoding region of human carcinoembrionic
 ID
     Q04592 standard; DNA; 324 BP.
 AC
     004592;
     28-SEP-1990 (first entry)
 DE
     Sequence encoding region of human carcinoembrionic light chain
 DE
     antigen (CEA).
 K₩
     Carcinoembryonic antiqen; CEA; Chelate-specific antiqen; CHA;
 KW
     tumor; cancer; serum sickness.
 OS
     Homo sapiens.
 PN
     EP-369566-A.
 PD
     23-MAY-1990.
 PF
     8-MAR-1989; 302313.
 PR
     17-NOV-1988; US-274105; EP-302313.
 PA
     (HYBR-) Hubritech Inc.
 PΙ
     Johnson MJ, Phelps JL;
 DR
     WPI; 90-157695/21.
 DR
     P-PSDB; R04936.
 PT
     Bifunctional chimeric antibodies -
 PT
     having variable regions which recognise different antigens and
PT
     metal chelates and human constant regions.
PS
     Claim 2; Page 24; 40pp; English.
CC
     Gene encodes portion of CEA light chain antigen, useful in
 CC
     generating chimeric monoclonal antibody binding to CEA at a tumour
CC
     site and a metal chelate bonded to say a toxin or other drug.
 SQ
     Sequence
               324 BP;
                       85 A;
                               83 C;
                                    77 G;
```

Conservative Substitutions

6aps

```
Initial Score
                   256
                        Optimized Score =
                                            257 Significance = 18.60
Residue Identity =
                   80%
                        Matches
                                            257 Mismatches
Gaps
                     0
                        Conservative Substitutions
                                                                 0
          10
                   20
                            30
                                     40
                                              50
                                                                70
                                                       60
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      1111 111111111111 | 111 1111111111
   GACATTGTGATGACCCAGTCTCAAAAATTCATGTCCACATCAGTAGGAGACAGGGTCAGCATCACCTGCAAG
   X
          10
                   20
                            30
                                     40
                                              50
         80
                  90
                          100
                                   110
                                            120
                                                    130
                                                              140
   GCCAGTCAGAGTGTGACTAATGATGTAGCTTGGTACCAACAGAAGCCAGGGCAGTCTCCTAAACTGCTGATA
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                  90
                         100
                                  110
                                           120
                                                    130
                                                             140
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               160
                        170
                                 180
                                          190
                                                   200
                                                            210
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      150
               160
                        170
                                 180
                                          190
                                                   200
                                                           210
    220
             230
                      240
                               250
                                        260
                                                 270
                                                          280
   TTCACCATCAGCACTGTGCAGGCTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTAC
    220
             230
                      240
                               250
                                        260
                                                 270
                                                          280
  290
           300
                    310
   ACGTTCGGAGGGGGGACCAAGCTGGAGATC
   ACGTTCGGTGCTGGGACCAAGCTGGAGCTGAAACGG
  290
           300
                    310
                             320
7. US-07-977-702-2 (1-318)
  N90813
              DNA encoding light chain variable region against c
 ID
     N90813 standard; cDNA; 324 BP.
AC
     N90813;
 DT
     24-JAN-1990. (first entry)
DE
     DNA encoding light chain variable region against carcinoembryonic antigen
     Carcinoembyonic antigen; immunogen; tumour; chimeric antibody.
KW
     Murine (mouse) hybridoma CEM 231.6.7 (ATCC HB 9620).
OS
FH
                   Location/Qualifiers
     Key
FT
     CDS
                   1..324
FT
     /*tag= a
PN
     EP-332424-A.
PD
     13-SEP-1989.
PF
     08-MAR-1989; 302312.
PR
     09-MAR-1988; US-272577.
PA
     (HYBR) Hybritech Inc.
PΙ
     Beidler CB, Johnson MJ, Ludwig JR, Carlo DJ, David GS;
DR
     WPI; 89-265635/37.
DR
     P-PSDB; P91381.
PT
     DNA constructs encoding light or heavy chain variable regions
PT
     - chimeric monoclonal antibody, specific for human
PT
     carcinoembryonic antigen, useful for diagnosis, imaging and
PT
     treatment of tumours.
PS
     Claim 2; page 18; 32pp; English.
CC
     This region contained in a DNA construct with heavy chain variable
CC
     and constant regions, light chain constant region, and 2 eukaryotic
CC
     leader sequences. The chimeric monoclonal antibodies thus formed
CC
     can be directed against CEA for immunoassay, and imaging and
```

treatment of tumours, eq colorectal or breast carcinoma. The

CC

```
CC
     than entirely murine Abs.
     Sequence
                        85 A; 83 C; 77 G;
 SQ
               324 BP;
Initial Score
                   256 Optimized Score = 257 Significance = 18.60
Residue Identity =
                   80% Matches
                                           257 Mismatches
Gaps
                     O Conservative Substitutions
                                                                 0
          10
                   20
                            30
                                     40
                                              50
                                                       60
                                                                70
   AGTATTGTGATGACCCAGACTCCCAAATTCCTGCTTGTTTCAGCAGGAGACAGGGTTACCATAACCTGCAAG
      1111111111111111 111 1111111 11
                                      1111 111111111111 1 111 1111111111
   GACATTGTGATGACCCAGTCTCAAAAATTCATGTCCACATCAGTAGGAGACAGGGTCAGCATCACCTGCAAG
   X
          10
                   20
                            30
                                     40
                                              50
                                                       60
                                                                70
         80
                 90
                         100
                                  110
                                           120
                                                    130
                                                             140
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                  GCCAGTCAGAATGTTCGTACTGCTGTTGCCTGGTATCAACAGAAACCAGGGCAGTCTCCTAAAGCACTGATT
         80
                 90
                         100
                                  110
                                           120
                                                    130
      150
               160
                        170
                                 180
                                         190
                                                  200
                                                           210
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   TACTTGGCATCCAACCGGTACACTGGAGTCCCTGATCGCTTCACAGGCAGTGGATCTGGGACAGATTTCACT
      150
                        170
               160
                                 180
                                         190
                                                  200
    220
             230
                               250
                      240
                                        260
                                                 270
                                                         280
   TTCACCATCAGCACTGTGCAGGCTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTAC
    220
             230
                      240
                               250
                                        260
                                                 270
                                                         280
  290
           300
                    310
                             X
   ACGTTCGGAGGGGGGACCAAGCTGGAGATC
   ACGTTCGGTGCTGGGACCAAGCTGGAGCTGAAACGG
  290
           300
                    310
                             320
8. US-07-977-702-2 (1-318)
  027132
              Light chain variable region contg. onpA leader pep
ID
     027132 standard; DNA; 387 BP.
AC
     027132;
DT
     21-JAN-1993 (first entry)
     Light chain variable region contg. ompA leader peptide.
DE
KW
     Heavy; antibody; Fv; CEM231.6.7; VL; ss; chelator.
05
     Synthetic.
     EP-497585-A.
PN
PD
     05-AUG-1992.
PF
     30-JAN-1992; 300775.
PR
     30-JAN-1991; US-647901.
PA
     (ANDE/) ANDERSON L D.
     (COOK/) COOK J A.
PA
PA
     (DAVI/) DAVID G S.
PA
     (HOCH/) HOCHSCHWENDER S M.
PA
     (KASH/) KASHER M S.
PA
     (SMIT/) SMITH M C.
PA
     (STEM/) STEMMER ₩ P C.
PI
     Anderson LD, Cook JA, David GS, Hochschwenderder SM, Kasher MS;
PΙ
     Smith MC, Stemmer WPC;
DR
     WPI; 92-261005/32.
PT
     Immobilisation and purification of molecules - using kinetically
PT
     inert immobilised metal-CP-protein complex useful in assau
PT
PS
```

Example 5; Page 37; 88pp; English.

constant regions are numan-derived and so are less immunogenic

```
Sequencing of cloned CEM231.6./. heavy and light chain variable
 CC
     regions along with the His-Trp-His-His-His fusion peptide was
 CC
     performed by standard procedures. This sequence shows the light
 CC
     chain variable sequence contg. the ompA leader peptide.
 CC
     See also @27108-33.
 SQ
     Sequence
               387 BP;
                         100 A;
                                   100 C;
                                            96 G;
                                                    91 T;
                    256 Optimized Score =
Initial Score
                                             257
                                                 Significance = 18.60
Residue Identity =
                    80% Matches
                                             257
                                                 Mismatches
Gaps
                      0
                        Conservative Substitutions
                                                                   0
                                                         10
                                                                  20
                                                 AGTATTGTGATGACCCAGACTC
                                                    111111111111111 141
   CTATCGCGATCGCAGTGGCACTGGCTGGTTTCGCCACCGTGGCGCAGGCCGACATTGTGATGACCCAGTCTC
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                 30
                          40
                                             60
                                                      70
         30
                  40
                           50
                                              70
                                                       80
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                                                                90
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                   AAAAATTCATGTCCACATCAGTAGGAGACAGGGTCAGCATCACCTGCAAGGCCAGTCAGAATGTTCGTACTG
      90
              100
                        110
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                                                            150
      100
               110
                        120
                                           140
                                  130
                                                    150
                                                             160
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    11111 11 1 11111111 11 1111
   CTGTTGCCTGGTATCAACAGAAACCAGGGCAGTCTCCTAAAGCACTGATTTACTTGGCATCCAACCGGTACA
   160
            170
                      180
                               190
                                        200
                                                 210
                                                          220
    170
                                200
             180
                       190
                                         210
                                                  220
                                                           230
   CTGGAGTCCCTGATCGCTTCACTGGCAGTGGATATGGGACGGATTTCACTTTCACCATCAGCACTGTGCAGG
   CTGGAGTCCCTGATCGCTTCACAGGCAGTGGATCTGGGACAGATTTCACTCTCACCATTACCAATGTGCAAT
 230
           240
                    250
                             260
                                      270
                                               280
                                                         290
                                                                  300
  240
            250
                     260
                              270
                                       280
                                                290
                                                          300
                                                                   310
   CTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTACACGTTCGGAGGGGGGACCAAGC
   CTGAAGACCTGGCAGATTATTTCTGTCTGCAACATTGGAATTATCCGCTCACGTTCGGTGCTGGGACCAAGC
         310
                  320
                           330
                                    340
                                              350
                                                       360
                                                                370
          X
   TGGAGATC
   HHHH
   TGGAGCTGAAATAG
       380
9. US-07-977-702-2 (1-318)
  928256
              Fv heavy chain/light chain variable domain fusion
ID
     Q28256 standard; DNA; 748 BP.
AC
     028256;
DT
     11-FEB-1993 (first entry)
     Fv heavy chain/light chain variable domain fusion protein.
DE
KW
     Minigene; pMZ18/1; monoclonal antibody; kappa; light chain;
KW
     variable region; pWW31; tuomour; ss.
OS
     Mus ausculus.
FH
     Key
                    Location/Qualifiers
FT
     CDS
                    5..731
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FT
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     /≇taq= b
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FT
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     /≇tag= d
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     EP-502812-A.
 PD
     09-SEP-1992.
 PF
     27-JAN-1992; 810056.
 PR
     05-FEB-1991; EP-810079.
 PA
     (CIBA ) CIBA GEIGY AG.
 PΙ
     Groner B. Hardman N. Harwerth I. Hynes NE. Wels WS;
 PI
     Zwickl M;
     WPI; 92-302096/37.
 DR
     P-PSDB; R26979.
 DR
 PT
     Recombinant antibodies directed to growth factor receptor C-erbB-2 -
 PT
     for diagnosing and treating tumours expressing C-erbB-2 e.g. breast
 PT
     or ovarian tumours
 PS
     Disclosure; Page 31-33; 67pp; English.
 CC
     The sequence given encodes the Fv heavy chain/light chain variable
 CC
     domain fusion protein which binds to the extracellular domain of the
 CC
     growth factor receptor c-erbB-2. This recombinant antibody can be used
 CC
     for the qualitative and quantitative determination of c-erbB-2. This
 CC
     can be used for monitoring or in-vivo localisation of tumours
 CC
     overexpressing c-erbB-2.
 SQ
     Sequence 748 BP;
                          178 A;
                                   190 C;
                                             192 G;
                                                      188 T;
Initial Score
                    256 Optimized Score =
                                             259 Significance = 18.60
Residue Identity =
                    81% Matches
                                             259 Mismatches
                                                                   59
Gaps
                      O Conservative Substitutions
                                                                    0
                                                  X
                                                         10
                                                                   20
                                                  AGTATTGTGATGACCCAGACTC
                                                     Ш
                                                          1 1111111 111
   370
                   380
                            390
                                      400
                                               410
                                                         420
                                                                  430
                  40
         30
                            50
                                     60
                                              70
                                                        80
   CCAAATTCCTGCTTGTTTCAGCAGGAGACAGGGTTACCATAACCTGCAAGGCCAGTCAGAGTGTGACTAATG
                  ACAAATTCCTGTCCACTTCAGTAGGAGACAGGGTCAGCATCACCTGCAAGGCCAGTCAGGATGTGTATAATG
        440
                 450
                           460
                                    470
                                             480
                                                       490
                                                                500
      100
                         120
                110
                                  130
                                            140
                                                     150
   ATGTAGCTTGGTACCAACAGAAGCCAGGGCAGTCTCCTAAACTGCTGATATATTATGCATCCAATCGCTACA
    CTGTTGCCTGGTATCAACAGAAACCAGGACAATCTCCTAAACTTCTGATTTACTCGGCATCCTCCCGGTACA
      510
               520
                         530
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                                            550
                                                     560
                                                              570
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              180
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                                200
                                          210
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                                                            230
```

Гί

I I ADEI - CAVIU

```
CIGGAGICCCIGAICGCIICACIGGCAGIGGAIAIGGGACGGAIIICACIIICACIAICAGCACIGIGCACIGIGCAG
    CTGGAGTCCCTTCTCGCTTCACTGGCAGTGGCTCTGGGCCGGATTTCACTTTCACCATCAGCAGTGTGCAGG
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              590
                       600
                                610
                                          620
                                                   630
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                     260
                               270
                                        280
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                                                           300
                                                                    310
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    CTGAAGACCTGGCAGTTTATTTCTGTCAGCAACATTTTCGTACTCCATTCACGTTCGGCTCGGGGACAAAAT
   650
            660
                     670
                               680
                                        690
                                                 700
                                                           710
          X
   TGGAGATC
    1 111111
   TCGAGATCTAGCTGATCAAAGCTCTAGA
          730
                   740
10. US-07-977-702-2 (1-318)
                Sequence encoding variable region of murine AHT 54
 ID
     905557 standard; DNA; 411 BP.
 AC
     005557;
 DT
     10-DEC-1990 (first entry)
     Sequence encoding variable region of murine AHT 54 light chain.
 DE
 KW
     Interleukin-2 receptor; IL-2; tumour necrosis factor; TNF; ss;
 05
FH
     Keu
                    Location/Qualifiers
 FT
     CDS
                    58..411
 FT
     /*tag= a
 PN
     EP-380068-A.
PD
     01-AUG-1990.
 PF
     24-JAN-1990; 101351.
 PR
     24-JAN-1989; US-301216.
PR
     04-DEC-1989; US-441702.
PA
     (MOLE-) MOLECULAR THERAPEU.
PI
     Zerler B;
 DR
     WPI; 90-232892/31.
 DR
     P-PSDB; R06253.
     Expression vectors for producing chimeric monoclonal antibodies -
PT
PT
     which express human constant region and non-human variable region
PS
     Disclosure; p; English.
 CC
     MAbs comprising mouse CH and CL constant regions whith human
 CC
     variable regions may be used to create mouse/human hybrid MAbs.
 CC
     which have a longer serum half-life. Method can be used to produce
CC
     Abs against interleukin-2 receptor and tumour necrosis factor.
 SQ
     Sequence
             411 BP;
                          110 A;
                                   94 C;
                                            104 G;
Initial Score
               =
                    255 Optimized Score =
                                              259 Significance = 18.52
Residue Identity =
                                              260 Mismatches
                    81% Matches
                                                                   58
Gaps
                      1 Conservative Substitutions
                                                                    0
                                                           10
                                                  AG-TATTGTGATGACCCAGACT
                                                  11 11111111111111111111
   CAGACTCAGGTCTTTGTATACATGTTGCTGTGGTTGTTCTGGTGGTGATGGAGACATTGTGATGACCCAGTCT
     40
               50
                        60
                                 70
                                           80
                                                  X 90
                                                             100
                   40
                             50
                                      60
                                               70
                                                         80
                                                                  90
   CCCAAATTCCTGCTTGTTTCAGCAGGAGACAGGGTTACCATAACCTGCAAGGCCAGTCAGAGTGTGACTAAT
                    CAAAAATTCATGTCCACATCAGTAGGAGACAGGGTCAGCGTCACCTGCAAGGCCAGTCAGAATGTGGGTACT
  110
            120
                     130
                              140
                                        150
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       100
                110
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                                   130
                                             140
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```

GATGTAGCTTGGTACCAACAGAAGCCAGGGCAGTCTCCTAAACTGCTGATATATTATGCATCCAATCGCTAC

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AATGTAGCCTGGTATCAACAGAAACCAGGGCAATCTCCTAAAACACTGATTTACTCGGCATCCTACCGTTAC
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                   200
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              180
                       190
                                 200
                                          210
                                                   220
                                                            230
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   <u>AGTGGAGTCCCTGATCGCTTCACAGGCAGTGGATCTAGGACAGATTTCACTCTCACCATCAGCAATGTGCAG</u>
                 270
                          280
                                   290
                                             300
                                                      310
                                                               320
   240
            250
                      260
                               270
                                        280
                                                 290
                                                           300
   GCTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTACACGTTCGGAGGGGGGACCAAG
    TCTGAAGACTTGGCAGAGTATTTCTGTCAGCAATATAACAGCTATCCTTGGACGTTCGGTGGAGGCACCAAG
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               340
                        350
                                  360
                                           370
                                                    380
                                                             390
 310
           X
   CTGGAGATC
   11111 111
   CTGGAAATCAAACGG
    400
          X 410
11. US-07-977-702-2 (1-318)
   N40022
               cDNA insert of pK17G4 encoding kappa anti-carcinoe
 ID
     N40022 standard; cDNA; 882 BP.
AC
     N40022;
DT
     01-DEC-1991 (first entry)
DE
     cDNA insert of pK17G4 encoding kappa anti-carcinoembryonic antigen.
K₩
     Immunoglobulin; ds DNA; carcinoembryonic antigen; vector pK17G4.
PN
     EP-125023-A.
PD
     14-NOV-1984.
PF
     06-APR-1984; 302368.
PR
     08-APR-1983; US-483457.
PA
     (CITY) City of Hope.
PA
     (GETH) Genentech Inc.
ΡI
     Cabilly S, Holmes WE, Wetzel RB, Heyneker HL, Riggs AD.
DR
     WPI; 84-283749/46.
PT
     Immunoglobulin(s) produced by recombinant host cells - useful as
PT
     antibodies analogous to forms from mammals.
PS
     Disclosure; Fig. 2A-B; 79pp; English.
CC
     The cDNA is contained within recombinant vector pK17G4 and encodes
CC
     kappa anti-carcinoembryonic antigen chain. Using the vector the
CC
     immunoglobulin is produced readily in pure monoclonal form. Genetic
CC
     manipulations can be used to produce chimeras of variants drawing
CC
     their homology from species differing from each other. Protein
CC
     manipulation is also possible.
SQ
     Sequence
              882 BP;
                         231 A;
                                  243 C;
                                            194 G:
                                                     214 T;
Initial Score
                    252 Optimized Score =
                                             254 Significance = 18.29
Residue Identity =
                    79% Matches
                                             255 Mismatches
                                                                  63
Gaps
                      1 Conservative Substitutions
                                                                   0
                                    10
                                              20
                                                       30
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                               GTTGCTGTGGTTGTCTGATGAAGGAGACATTGTGATGACCCAGTCTCACAAATTCATGTCCACATCAGT
           10
                    20
                             30
                                      40
                                               50
                                                         60
                                                                  70
                60
                         70
       50
                                   80
                                            90
                                                    100
   AGGAGACAGGGTTACCATAACCTGCAAGGCCAGTCAGAGTGTGACTAATGATGTAGCTTGGTACCAACAGAA
   AGGAGACAGGGTCAGCATCACCTGCAAGGCCAGTCAGGATGTGGGTGCTGCTATAGCCTGGTATCAACAGAA
         80
                  90
                          100
                                             120
                                   110
                                                      130
                                                               140
```

11111 11 1

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     ACCAGGACAATCTCCTAAACTACTGATTTACTGGGCATCCACCCGGCACACTGGAGTCCCTGATCGCTTCAC
      150
                160
                          170
                                             190
                                                      200
                                                                210
   190
            200
                      210
                               220
                                         230
                                                  240
                                                            250
                                                                      260
    TGGCAGTGGATATGGGACGGATTTCACTTTCACCATCAGCACTGTGCAGGCTGAAGACCTGGCAGTTTATTT
     AGGCAGTGGATCTGGGACAGATTTCACTCTCACCATTAGCAATGTGCAGTCTGATGACTTGGCAGATTATTT
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              230
                                                    270
          270
                    280
                             290
                                       300
                                                 310
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                    1 1 111
                             111111111 | 111111111111111 |
    CTGTCAACAATATAGCGGGTATCCTCTCACGTTCGGTGCTGGGACCAAGCTGGAGCTGAAACGGGCTGATGC
   290
            300
                      310
                               320
                                         330
                                                  340
                                                          X 350
                                                                     360
    TGCACCAACTGTATCCATCTTCCCACCATCCAGTGA
          370
                    380
                             390
12. US-07-977-702-2 (1-318)
    004264
                Encodes Colon Cancer monoclonal antibody CC83 ligh
 ID
     Q04264 standard; DNA; 685 BP.
 AC
     Q04264;
     13-SEP-1990 (first entry)
 DT
 DE
     Encodes Colon Cancer monoclonal antibody CC83 light chain variable region
 KW
     chimeric antibodies; TAG72; light chain variable region;
 KW
     heavy chain variable region; ss
 05
      synthetic.
 FH
     Key
                     Location/Qualifiers
 FT
      intron
                     136..313
 FT
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 FT
     transit_peptide 76..325
 FT
     /*tag= b
 FT
     /note="intron is spliced out"
 FT
     misc feature
                     326..610
 FT
     /#tag= c
 FT
     /label= encodes variable segment
 FT
     misc_feature
                    611..646
 FT
     /*tag= d
 FT
     /label=encodes J4 segment
 PN
     EP-365997-A.
 PD
     02-MAY-1990.
 PF
     18-OCT-1989; 119361.
 PR
     19-0CT-1988;US-259943.
      (DOWC) Dow Chemical Co.
 PΙ
     Mezes P. Gourlie B. Rixon M;
 DR
     WPI; 90-133521/18.
 DR
     P-PSDB; R04387.
 PT
     Chimeric antibodies against TAG72 and conjugate to provide imaging
 PT
     markers and therapeutic tools
 PS
     Disclosure; pp; English.
     The polypeptide encoded by this sequence forms part of a chimera. The
 CC
 CC
     other components are a heavy chain variable region and human-derived
 CC
     constant light and heavy chain regions. The variable regions have
 CC
     high affinity for TAG72. The constant regions reduce the side-effects
 CC
     when administered to human patients because they are of human origin.
 CC
      See also 004258-63 and 004265-9.
 50
      Sequence
                685 BP; 187 A; 138 C;
Initial Score
                     250 Optimized Score =
                                              253 Significance = 18.13
Residue Identity =
                     79% Matches
                                               254 Mismatches
Gaps
                       1
                         Conservative Substitutions
                                                                =
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```

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      280
               290
                        300
                                 310
                                          320
                                                X 330
                                                            340
         30
                   40
                            50
                                     60
                                              70
                                                       80
   CCCAAATTCCTGCTTGTTTCAGCAGGAGACAGGGTTACCATAACCTGCAAGGCCAGTCAGAGTGTGACTAAT
                350
             360
                      370
                               380
                                         390
                                                 400
       100
                110
                         120
                                  130
                                           140
                                                    150
                                                             160
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   GCTGTAGCCTGGTTTCAGCATAAACCAGGTCAGTCTCCTAAACTACTGATTTACGGGGCATCCACCCGGCAC
  420
           430
                    440
                             450
                                       460
                                                470
                                                         480
                                                                  490
     170
              180
                       190
                                200
                                         210
                                                  220
                                                           230
   ACTGGAGTCCCTGATCGCTTCACTGGCAGTGGATATGGGACGGATTTCACTTTCACCATCAGCACTGTGCAG
   ACTGGAGTCCCTGATCGCTTCACAGGCAGTGGATCTGGGACAGATTTCACTCTCATCATTAGCAATGTGCAG
         500
                   510
                            520
                                     530
                                              540
                                                       550
                                                                560
   240
            250
                                        280
                     260
                               270
                                                 290
                                                          300
   GCTGAAGACCTGGCAGTTTATTTCTGTCAGCAGGATTATAGCTCTCCGTACACGTTCGGAGGGGGGACCAAG
    1111 111 111111 111111 111111 11
                                        111 111 1 1111111
   TCTGAGGACTTGGCAGATTATTTGTGTCAGCATTATAGCGGCTATCCATTCACGTTCGGCTCGGGGACAAAG
        570
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                          590
                                   600
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 310
          X
   CTGGAGATC
    1111 11
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      640 X
               650
                        660
                                 670
                                          680
13. US-07-977-702-2 (1-318)
   N40023
               mRNA encoding kappa anti-carcinoembryonic antigen.
     N40023 standard; mRNA; 882 BP.
ID
AC
     N40023;
DT
     01-DEC-1991 (first entry)
     mRNA encoding kappa anti-carcinoembryonic antigen.
K₩
     Immunoglobulin; ss mRNA; carcinoembryonic antigen.
FH
     Key
                   Location/Qualifiers
FT
     CDS
                   1..673
FT
     /#taq= a
PN
     EP-125023-A.
     14-NOV-1984.
PD
PF
     06-APR-1984; 302368.
PR
     08-APR-1983; US-483457.
PA
     (CITY) City of Hope.
PA
     (GETH) Genentech Inc.
PI
     Cabilly S. Holmes WE, Wetzel RB, Heyneker HL, Riggs AD.
DR
     WPI; 84-283749/46.
DR
     P-PSDB; P40031.
     Immunoglobulin(s) produced by recombinant host cells - useful as
PT
PT
     antibodies analogous to forms from mammals.
PS
     Disclosure; Fig. 3; 79pp; English.
CC
     The ARNA is the coding sequence of the cDNA fragment contained within
CC
     recombinant vector pK17G4 (see N40022). It encodes kappa anti-
CC
     carcinoembryonic antigen chain. Using the vector the immunoglobulin is
CC
     produced readily in pure monoclonal form. Genetic manipulations can be
```

used to produce chimeras of variants drawing their homology from species

CC

X

10

```
50
     Sequence
              882 BP;
                         231 A;
                                  244 C;
                                           194 G;
Initial Score
                   249 Optimized Score =
                                            253 Significance = 18.06
Residue Identity =
                   59% Matches
                                            254 Mismatches
                                                                 64
Gaps
                     1 Conservative Substitutions
                                                                  0
                            X
                                    10
                                             20
                                                      30
                                                               40
                            AG-TATTGTGATGACCCAGACTCCCAAATTCCTGCTTGTTTCAGC
                            GUUGCUGUGGUUGUCUGGUGUUGAAGGAGACAUUGUGAUGACCCAGUCUCACAAAUUCAUGUCCACAUCAGU
          10
                   20
                                      40
                                               50
       50
                60
                         70
                                           90
                                                   100
                                                            110
                                  80
   AGGAGACAGGGTTACCATAACCTGCAAGGCCAGTCAGAGTGTGACTAATGATGTAGCTTGGTACCAACAGAA
   AGGAGACAGGGUCAGCAUCACCUGCAAGGCCAGUCAGGAUGUGGGUGCUGCUAUAGCCUGGUAUCAACAGAA
                          100
                                   110
                                            120
                                                     130
    120
             130
                      140
                               150
                                        160
                                                  170
                                                          180
   GCCAGGGCAGTCTCCTAAACTGCTGATATATTATGCATCCAATCGCTACACTGGAGTCCCTGATCGCTTCAC
    ACCAGGACAAUCUCCUAAACUACUGAUUUACUGGGCAUCCACCCGGCACACUGGAGUCCCUGAUCGCUUCAC
      150
               160
                        170
                                 180
                                          190
                                                   200
                                                            210
  190
           200
                    210
                             220
                                       230
                                                240
                                                         250
                                                                  260
   TGGCAGTGGATATGGGACGGATTTCACTTTCACCATCAGCACTGTGCAGGCTGAAGACCTGGCAGTTTATTT
    AGGCAGUGGAUCUGGGACAGAUUUCACUCUCACCAUUAGCAAUGUGCAGUCUGAUGACUUGGCAGAUUAUUU
    220
             230
                      240
                               250
                                        260
                                                 270
         270
                   280
                            290
                                     300
                                              310
                                                       X
   CTGTCAGCAGGATTATAGCTCTCCGTACACGTTCGGAGGGGGGGACCAAGCTGGAGATC
                   1 | 111
                            11111111 | 111111111111 | 1
   CUGUCAACAAUAUAGCGGGUAUCCUCUCACGUUCGGUGCUGGGACCAAGCUGCAGCUGAAACGGGCUGAUGC
  290
           300
                    310
                             320
                                       330
                                                340
                                                       X 350
   UGCACCAACUGUAUCCAUCUUCCCACCAGCCAGUGA
          370
                   380
                            390
14. US-07-977-702-2 (1-318)
   025592
               Encodes 4D5 Fab IgG antibody - recognises HER-2 re
 ID
     Q25592 standard; DNA; 2178 BP.
 AC
     025592;
 DT
     08-DEC-1992 (first entry)
 DE
     Encodes 4D5 Fab IgG antibody - recognises HER-2 receptor.
 KW
     humanised IgG antibody; human growth hormone; hGH; selection;
 KW
     screening; ss.
 05
     Homo sapiens.
FH
     Key
                   Location/Qualifiers
FT
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     exon
FT
     /*tag= a
FT
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FT
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                   796..2178
FT
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PN
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 PD
     11-JUN-1992.
 PF
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PR
     03-DEC-1990; US-621667.
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     10-APR-1991; US-683400.
PR
     14-JUN-1991; US-715300.
 PR
     08-AUG-1991; US-743614.
 PA
     (GETH ) GENENTECH INC.
```

differing from each other. Protein manipulation is also possible.

```
PΙ
          Matthews DJ, Wells JA;
  DR
          WPI; 92-217069/26.
  DR
          P-PSDB; R24047,8.
  PT
          Selecting and enriching variant proteins - comprises fusing gene
  PT
          encoding e.g. growth hormone to part of MI3 phage coat protein
  PT
           and autagenising fusion prior to selection
  PS
          Claim 46; Fig 11; 102pp; English.
  CC
          This sequence encodes the Fab portion of the 4D5 humanised IqG
  CC
           antibody that recognises the HER-2 receptor. It was inserted into
  CC
          the plasmid pS0132, which had the DNA encoding human growth hormone
  CC
           excised from it. The plasmid was used to transform E. coli SR101.
  CC
          The sequence encodes two genes. The first encodes the variable and
  CC
          constant regions of the light chain, and with the st II signal
  CC
           sequence at its 5' end. The second contains four regions - the st II
  CC
          signal sequence at its 5' end, the variable domain of the heavy chain,
  CC
          the first domain of the heavy chain constant region, followed by the
  CC
           Mi3 gene III.
 SQ
          Sequence
                             2178 BP;
                                                   498 A;
                                                                     572 C;
                                                                                      549 G;
                                                                                                        559 T;
Initial Score
                             =
                                      234 Optimized Score =
                                                                                      237 Significance = 16.89
Residue Identity =
                                      74% Matches
                                                                                      237 Mismatches
                                                                                                                               81
                                          O Conservative Substitutions
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                                                                                                   IIII
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     20
                       30
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                                                          50
                                                                           60
                                                                                              70
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                                                                                                                                 90
                  30
                                                     50
                                                                      60
                                                                                        70
                                                                                                          80
                                                                                                                           90
       CCAAATTCCTGCTTGTTTCAGCAGGAGACAGGGTTACCATAACCTGCAAGGCCAGTCAGAGTGTGACTAATG
       1 1 | 1111 | 1 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 11 | 
                                                                                               111111111 11111 11 11
       CGAGCTCCCTGTCCGCCTCTGTGGGCGATAGGGTCACCATCACCTGCCGTGCCAGTCAGGATGTGAATACTG
                                   110
                                                    120
                                                                      130
                                                                                        140
                                                                                                          150
                                                                                                                           160
            100
                              110
                                               120
                                                                 130
                                                                                   140
                                                                                                    150
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       ATGTAGCTTGGTACCAACAGAAGCCAGGGCAGTCTCCTAAACTGCTGATATATTATGCATCCAATCGCTACA
         170
                                180
                                                 190
                                                                   500
                                                                                    210
                                                                                                      220
                                                                                                                        230
        170
                          180
                                            190
                                                             200
                                                                               210
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                                                                                                                  230
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                                                                                                               300
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       1111
                                                                                              111111111 111 111111
       CGGAAGACTTCGCAACTTATTACTGTCAGCAACATTATACTACTCCTCCCACGTTCGGACAGGGTACCAAGG
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                        320
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                                                            340
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                                                                                               360
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       11111111
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                                                        410
                                                                          420
                                                                                            430
```

pass as darrard to, breene k, menner by, Lowman MB;

15. US-07-977-702-2 (1-318)
Q28747 Anti-CEA specific light chain variable region DNA.

ID @28747 standard; DNA; 387 BP.

AC 928747;

Гі

```
VI-MAK-1993 (first entry)
 DE
     Anti-CEA specific light chain variable region DNA.
 KW
     Human; carcinoembryonic antigen; heavy chain; light chain; variable;
 KW
     region; diagnostic; tumour; markers; targetting; ss.
 08
     Mus Ausculus.
 FH
                    Location/Qualifiers
     Keu
 FT
     sig_peptide
                    1..60
 FT
     /#taq= a
 FT
     mat_peptide
                    61..387
 FT
     /*taq= b
 PN
     J04234987-A.
 PD
     24-AUG-1992.
 PF
     28-DEC-1990; 408810.
 PR
     28-DEC-1990; JP-408810.
 PA
     (MITU ) MITSUBISHI KASEI CORP.
 DR
     WPI; 92-327631/40.
 DR
     P-PSDB; R27054.
 PT
     New DNA fragments encoding variable regions of ABS specific for
 PT
     human CEA - for diagnosing and monitoring tumours, as tumour
 PT
     markers and for treatment of tumours
 PS
     Claim 1; Fig 2; 7pp; Japanese.
 CC
     The anti-CEA murine monoclonal antibody light chain variable region
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     was obtd. by screening a cDNA library prepd. from mRNA obtd. from
 CC
     hybridomas producing anti-CEA-specific antibodies with a probe
 CC
     based on the constant region of the L-chain. The antibodies reacts
 CC
     specifically with human CEA and are useful as a diagnostic agents,
 CC
     as tumour markers for digestive organs, for diagnosis of malignant
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     tumours; for monitering after cancer operations, to follow up bloodless
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     therapy or as therapeutic agents in passive innune therapy and
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                           See also 928744-6.
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FastDB - Fast Pairwise Comparison of Sequences
Release 5.4
Results file us-07-977-702-2.res made by shears on Tue 15 Jun 93 11:04:41-PDT.
Query sequence being compared: US-07-977-702-2 (1-318)
Number of sequences searched:
                                            125798
Number of scores above cutoff:
                                              4066
      Results of the initial comparison of US-07-977-702-2 (1-318) with:
  Data bank : EMBL-NEW 3, all entries
  Data bank : GenBank 75, all entries
  Data bank : GenBank-NEW 3, all entries
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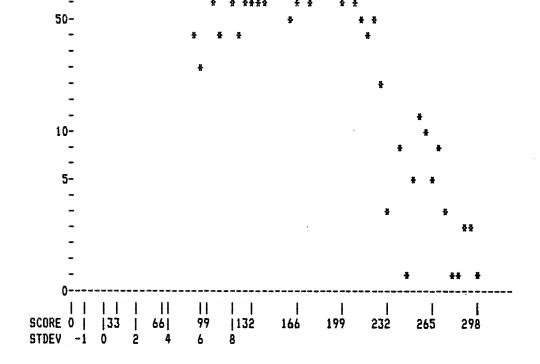
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370

M B Ε

U Ε N C Ε AACCTGGAAATAAAACGG



PARAMETERS

Similarity matrix	unitary	K-tuple	4
Mismatch penalty	1	Joining penalty	30
Gap penalty	1.00	Window size	32
Gap size penalty	0.33		
Cutoff score	0		
Randomization group	0		
Initial scores to save	40	Alignments to save	15
Optimized scores to sav	e 0	Display context	50

SEARCH STATISTICS

Scores:	Mean	Median	Standard Deviation		
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Times: CPU 00:28:23.05		Total Elapsed 00:58:37.00			

Number of residues: 150464018
Number of sequences searched: 125798
Number of scores above cutoff: 4066

Cut-off raised to 21.
Cut-off raised to 26.
Cut-off raised to 29.
Cut-off raised to 32.
Cut-off raised to 34.
Cut-off raised to 36.
Cut-off raised to 39.
Cut-off raised to 41.
Cut-off raised to 43.
Cut-off raised to 46.
Cut-off raised to 49.
Cut-off raised to 52.
Cut-off raised to 52.

The scores below are sorted by initial score. Significance is calculated based on initial score.

A 100% identical sequence to the query sequence was not found.

				Init.	Opt.				
Seque	nce Name	Description	Length			Sig.	Frame		
	**** 23 standard deviations above mean ****								
1.	MUSOHP57K	Mus musculus mRNA for Ig kapp		298	300	23.87	0		
2.	563304	Vk28 VkJk [not specified, Gen		295	297	23.61			
3.	MUSSCETK	Mus musculus DNA for Ig kappa			297	23.61			
4.	MMIGRF28	Murine MRL-RF28L mRNA for IgM			298	23.61			
5.	MUSIGKCPF	Mouse Ig active kappa-chain (938	290	293	23.17			
6.	MMIGKL	Mouse mRNA for Ig C.C58 M75 k	938	290	293	23.17			
		**** 22 standard deviations							
7.	560946	mAb BA.N4:4.57 V kappa region	300	287	289	22.91	0		
8.	MMVL105	Mouse mRNA for kappa-immunogl		287	290	22.91			
9.	MUSIGKVM	Mouse Ig germline kappa-chain		280	294	22.29			
10.	S60873	Ab2 kappa chain V region mA		278	279	22.11			
		**** 21 standard deviations		ean **	F #				
11.	MMNPCRLB6	M.musculus NPCRLB6 mRNA for i	321	273	275	21.67	0		
12.	MUSIGKCNK	Mouse Ig rearranged kappa-cha	456	272	275	21.58	0		
13.	MMNPCRLB1	M.musculus NPCRLB1 mRNA for i	321	271	273	21.50	0		
14.	MUSIGKVN	Mouse Ig germline kappa-chain	2002	270	284	21.41	0		
15.	MUSIGKCKP	Mouse Ig kappa-chain mRNA V-r	341	269	272	21.32	. 0		
16.	MUSIGLAFF	Mouse Ig kappa-chain mRNA V r	318	268	271	21.23	0		
17.	S51704	hybridoma antibody kappa-chai	313	266	266	21.06	0		
18.	MUSIGKAFO	Mouse Ig active kappa-chain m	313	266	266	21.06	0		
19.	MUSIGLAFD	Mouse Ig kappa-chain mRNA V r	315	266	266	21.06	0		
20.	MUSIGKAL	Mouse Ig active kappa gene: M	739	266	269	21.06	0		
		**** 20 standard deviations	above me	an **	#				
21.	MMVL3190	Mouse mRNA for kappa-immunogl	321	264	264	20.88	0		
	MUSIGKCSR	Mouse Ig active kappa-chain m	321	262	265	20.70	0		
	MUSIGKAAV	Mouse Ig rearranged kappa-cha	363	262	270	20.70	0		
24.	MUSIGKAQ	Mouse Ig kappa active gene: k	765	261	263	20.61	0		
25.	S51746	hybridoma antibody kappa-chai	276	260	261	20.53	0		
	MUSIGKAFV	Mouse Ig active kappa-chain m	275	259	260	20.44	0		
27.	MMBV1745L	M.musculus light chain variab	324	259	261	20.44	0		
	MUSIGKAJ	mouse ig kappa mrna from морс	943	259	262	20.44	0		
	MMIGK9	Messenger RNA for mouse immun	943	259	262	20.44	0		
30.	MUSIGKAJA	Mouse Ig kappa active mRNA fr	462	258	261	20.35			
	MUSNC41	Mus musculus NC41 mRNA, parti	324	257	260	20.26			
	MMIGKV	Mouse mRNA for immunoglobulin	462	257	259	20.26	0		
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	MUSIGKAED	Mouse Ig active kappa chain m	321	256	258	20.17			
	MUSIGLAA	Mouse rearranged light chain	354	256	260	20.17	0		
	HUMIGLIA	Human Ig light chain VJ regio		256	258	20.17	0		
37.	S115039	immunoglobulin light chain va	411	255	259	20.09	0		
		**** 19 standard deviations		ean **	F#				
	MUSIGKCNM	Mouse Ig active kappa chain m		254	259	20.00			
	MUSIGLAB	Mouse rearranged light chain	357	254	260	20.00	0		
40.	MUSIGLAFE	Mouse Ig kappa-chain mRNA V r	303	253	257	19.91	0		

1. US-07-977-702-2 (1-318)

MUSOHP57K Mus musculus mRNA for Ig kappa chain precursor.

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LOCUS
            MUSOHP57K
                          322 bp ss-mRNA
                                                               11-NOV-1992
DEFINITION Mus musculus mRNA for Ig kappa chain precursor.
ACCESSION
KEYWORDS
            Ig kappa chain; steroid specific antibody; variable region.
SOURCE
            Mus musculus (strain BALB/c) hybridoma cell line OHP57.G6.1 mRNA.
  ORGANISM
            Mus musculus
            Eukaryota; Animalia; Chordata; Mammalia; Rodentia; Muridae.
REFERENCE
            1 (bases 1 to 322)
  AUTHORS
            Sawada, J., Mizusawa, S., Terao, T., Naito, M. and Kurosawa, Y.
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HILLE
           Molecular characterization of monoclonal anti-steroid antibodies:
           primary structures of the variable regions of seven antibodies
           specific for 17alpha-hydroxyprogesterone or 11-deoxycortisol and
           their pH-reactivity profiles
   JOURNAL
           Mol. Immunol. 28, 1063-1072 (1991)
  STANDARD
           full automatic
 COMMENT
           Submitted (27-JUL-1992) to DDBJ by: Jun-ichi Sawada
           Dept. of Biochem. and Immunochem.
           National Institute of Hygienic Sciences
           1-18-1 Kamiyoga
           Setagaya, Tokyo 158
           Japan
           Phone: 03-3700-1141 x240
           Fax:
                  03-3707-6950.
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BASE COUNT
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Initial Score
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Residue Identity =
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Gaps
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2. US-07-977-702-2 (1-318)

S63304 Vk28 VkJk [not specified, Genomic, 322 nt]

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 KEYWORDS
 SOURCE
           not specified
  ORGANISM
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           Unclassified.
 REFERENCE
           1 (bases 1 to 322)
  AUTHORS
           Sawada.J., Mizusawa.S., Terao.T., Naito.M. and Kurosawa.Y.
  TITLE
           Molecular characterization of monoclonal anti-steroid antibodies:
           primary structures of the variable regions of seven antibodies
           specific for 17 alpha-hydroxyprogesterone or 11-deoxycortisol and
           their pH-reactivity profiles.
  JOURNAL
           Mol. Immunol. 28, 1063-1072 (1991)
  STANDARD
           full automatic
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           This sequence comes from Fig 5.
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                        322 bp ds-DNA
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           Mus musculus DNA for Ig kappa chain precursor.
 DEFINITION
 ACCESSION
           D12737
 KEYWORDS
            Ig kappa chain; steroid specific antibody; variable region.
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           Mus musculus (strain BALB/c) hybridoma cell line SCET.M8.1.1 DNA.
  DRGANISM
           Mus musculus
           Eukaryota; Animalia; Chordata; Mammalia; Rodentia; Muridae.
 REFERENCE
           1 (bases 1 to 322)
  AUTHORS
           Sawada, J., Mizusawa, S., Terao, T., Naito, M. and Kurosawa, Y.
           Molecular characterization of monoclonal anti-steroid antibodies:
  TITLE
           primary structures of the variable regions of seven antibodies
           specific for 17alpha-hydroxyprogesterone or 11-deoxycortisol and
           their pH-reactivity profiles
           Mol. Immunol. 28, 1063-1072 (1991)
  JOURNAL
  STANDARD
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BASE COUNT
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                87 a
                                        79 t
                                84 q
ORIGIN
                                            297 Significance = 23.61
Initial Score
                   295
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Residue Identity =
                    93%
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Gaps
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3. US-07-977-702-2 (1-318)

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                           398 bp
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                                                               04-AUG-1992
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             X14622
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             autoantibody; Ig kappa light chain; Ig light chain; IgM;
             immunoglobulin; variable region.
 SOURCE
             BOUSE
   ORGANISM Mus musculus
             Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
             Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
             1 (bases 1 to 398)
   AUTHORS
             Kofler, R.
   TITLE
             Direct Submission
   JOURNAL
             Submitted (07-MAR-1989) Kofler R., General and Experimental
             Pathology, University of Innsbruck, Fritz-Pregl-Str.3, A-6020
             Innsbruck, Austria.
  STANDARD
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             2 (bases 1 to 362)
   AUTHORS
             Kofler, R., Duchosal, M.A. and Dixon, F.J.
   TITLE
             Complexity, polymorphism, and connectivity of mouse V-kappa gene
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   JOURNAL
             Immunogenetics 29, 65-74 (1989)
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 BASE COUNT
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 ORIGIN
Initial Score
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Residue Identity =
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Gaps
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5. US-07-977-702-2 (1-318)
  MUSIGKCPF
              Mouse Ig active kappa-chain (V-J-C) mRNA from C.C5
LOCUS
           MUSIGKCPF
                        938 bp ss-ARNA
                                               ROD
                                                        15-SEP-1989
DEFINITION Mouse Ig active kappa-chain (V-J-C) mRNA from C.C58 m75 myeloma.
ACCESSION
KEYWORDS
           constant region; immunoglobulin light chain; immunoglobulin-kappa;
            joining region; processed gene; variable region.
 SOURCE
           Mouse C.C58 m75 myeloma cDNA to mRNA, clone p9.
  ORGANISM Mus musculus
           Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
           Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
REFERENCE
           1 (bases 1 to 938)
           Goldrick, M.M., Boyd, R.T., Ponath, P.D., Lou, S.Y. and Gottlieb, P.D.
  AUTHORS
  TITLE
           Molecular genetic analysis of the VkSer group associated with two
           mouse light chain genetic markers
  JOURNAL
           J. Exp. Med. 162, 713-728 (1985)
  STANDARD full automatic
FEATURES
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BASE COUNT ORIGIN 248 a 260 c 206 g 224 t

Initial Score = 290 Optimized Score = 293 Significance = 23.17
Residue Identity = 92% Matches = 293 Mismatches = 25
Gaps = 0 Conservative Substitutions = 0

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6. US-07-977-702-2 (1-318)

MMIGKL Mouse mRNA for Ig C.C58 M75 kappa light chain (VK

LOCUS MMIGKL 938 bp RNA ROD 07-MAY-1992
DEFINITION Mouse mRNA for Ig C.C58 M75 kappa light chain (VK Ser-group)
ACCESSION X02816 M12177

KEYWORDS constant region; Ig light chain; immunoglobulin; joining region;

signal peptide; variable region.

SOURCE mouse

ORGANISM Mus musculus

Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia; Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.

REFERENCE 1 (bases 1 to 938)

AUTHORS Goldrick, M.M., Boyd, R.T., Ponath, P.D., Lou, S.Y. and Gottlieb, P.D. TITLE Molecular genetic analysis of the VkSer group associated with two

mouse light chain genetic markers

JOURNAL J. Exp. Med. 162, 713-728 (1985)

STANDARD full automatic

COMMENT BALB/C mice germline V(K1) region see MMIG25, C(K) region see

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Data kindly reviewed (24-FEB-1986) by P.D. Gottlieb.
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7. US-07-977-702-2 (1-318)
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LOCUS
                         300 bp
 DEFINITION
           mAb BA.N4:4.57 V kappa region [mice, B cell hybridoma, BALB/c,
            aRNA, 300 ntl
            S60946
 ACCESSION
 KEYWORDS
SOURCE
            mice BALB/c B cell hybridoma
  ORGANISM
           Unclassified.
            Unclassified.
 REFERENCE
            1 (bases 1 to 300)
  AUTHORS
            Carlsson, L., Andersson, A. and Holmberg, D.
  TITLE
            Germ-line origin of functional idiotypic interactions:
            identification of two idiotypically connected, natural antibodies
            that are encoded by germ-line gene elements.
            Eur. J. Imaunol. 21, 2285-2288 (1991)
  JOURNAL
  STANDARD
            full automatic
 COMMENT
            This entry [NCBI gibbsq 60946] was created by the journal scanning
            component of NCBI/GenBank at the National Library of Medicine.
            This sequence comes from Fig. 2.
FEATURES
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                    region feature. Description: mAb BA.N4:4.57 V kappa
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Initial Score
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Residue Identity =
                    96% Matches
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Gaps
                       Conservative Substitutions
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10

20

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50

60

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  MMVL105
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 LOCUS
            MMVL105
                         321 bp
                                  RNA
                                                 ROD
                                                          04-AUG-1992
 DEFINITION
            Mouse mRNA for kappa-immunoglobulin light chain V-J region (cell
            line 105-2H)
            X51852
 ACCESSION
 KEYWORDS
            Ig kappa light chain; Ig light chain; immunoglobulin;
            joining region; variable region.
 SOURCE
            nouse
  DRGANISM
            Mus musculus
            Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
            Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
            1 (bases 1 to 321)
  AUTHORS
            Reininger, L.
  TITLE
            Direct Submission
   JOURNAL
            Submitted (16-FEB-1990) Reininger L., Dept of Pathology , Centre
            Medical Universitaire, 1 rue michel servet, CH-1211 Geneve 4,
            Switzerland.
  STANDARD
            full automatic
 REFERENCE
            2 (bases 1 to 321)
  AUTHORS
            Reininger, L., Shibata, T., Ozaki, S., Shirai, T., Jaton, J.C. and
            Izui,S.
  TITLE
            Variable region sequences of pathogenic anti-mouse red blood cell
            autoantibodies from autoimmune NZB mice.
  JOURNAL
            Eur. J. Immunol. 20, 771-777 (1990)
  STANDARD
            full automatic
 COMMENT
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Residue Identity =
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Gaps
                         Conservative Substitutions
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LOCUS
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                                                ROD
                                                         30-JUN-1987
DEFINITION Mouse Ig germline kappa-chain V-region gene V-Ser.
ACCESSION
KEYWORDS
           constant region; germline; immunoglobulin-kappa; variable region.
SOURCE
           Mouse (Balb/c) liver DNA, clone EM3.BALB-1.
  DRGANISM
           Mus musculus
           Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
           Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
REFERENCE
           1 (bases 1 to 1816)
           Boyd, R.T., Goldrick, M.M. and Gottlieb, P.D.
  AUTHORS
  TITLE
           Structural differences in a single gene encoding the V-k-Ser group
           of light chains explain the existence of two mouse light-chain
           genetic markers
           Proc. Natl. Acad. Sci. U.S.A. 83, 9134-9138 (1986)
  JOURNAL
  STANDARD
           full automatic
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10. US-07-977-702-2 (1-318)
   S60873
               Ab2 kappa chain V region | mAb F {V/J junction} [m
LOCUS
            S60873
                        300 bp
                                                ROD
                                 ARNA
                                                         08-JAN-1992
DEFINITION
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            mRNA Partial, 300 ntl
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ACCESSION

560873

/note="ig kappa-chain signal peptide-

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SOURCE
           mice MLR-lpr/lpr
   ORGANISM Unclassified.
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 REFERENCE
           1 (bases 1 to 300)
  AUTHORS
           Koizumi, T., Puccetti, A., Migliorini, P., Barrett, K.J. and
           Schwartz, R.S.
  TITLE
           Molecular heterogeneity of auto-anti-idiotypic antibodies in
           MLR-lpr/lpr mice.
  JOURNAL
           Eur. J. Immunol. 21, 2185-2193 (1991)
  STANDARD
           full automatic
 COMMENT
           This entry [NCBI gibbsq 60873] was created by the journal scanning
           component of NCBI/GenBank at the National Library of Medicine.
           This sequence comes from Fig.6.
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Initial Score
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Residue Identity =
                   93% Matches
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Gaps
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11. US-07-977-702-2 (1-318)
   MMNPCRLB6
               M.musculus NPCRLB6 mRNA for immunoglobulin light c
     MMNPCRLB6 standard; RNA; ROD; 321 BP.
 ID
XX
AC
     X70097; X51612;
XX
DT
     19-MAR-1993 (Rel. 35, Created)
```

CUMUNIS

TŒ

19-MAR-1993 (Rel. 35, Last updated, Version 1)

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DE
     M.musculus NPCRLB6 mRNA for immunoglobulin light chain, variable
 DE
     region
 XX
KW
     immunoglobulin light chain variable region.
 XX
05
     Mus musculus (mouse)
DC.
     Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
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     Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
XX
RN
     [1]
RP
     1-321
RA
     Izui S.;
RT
RL
     Submitted (05-FEB-1993) on tape to the EMBL Data Library by:
RL
     S. Izui, Dept. of Pathology, Centre Medical Universitaire, 1, Rue
RL
     Michel Servet, CH-1211, geneva 4, SWITZERLAND
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CC
     *source: tissue=lymphoid;
CC
     *source: cell_type=hybridoma;
CC
     *source: cell_line=CB6;
CC
     #source: is_germline=N;
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                                           275 Significance = 21.67
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                                           275 Mismatches
                                                               43
                     O Conservative Substitutions
Gaps
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XX

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12. US-07-977-702-2 (1-318)
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                Mouse Ig rearranged kappa-chain mRNA, clone AN12K.
 LOCUS
            MUSIGKCNK
                          456 bp ss-aRNA
                                                   ROD
                                                             15-MAR-1990
 DEFINITION Mouse Ig rearranged kappa-chain mRNA, clone AN12K.
 ACCESSION
            M19914 J03832
 KEYWORDS
            constant region; immunoglobulin light chain; immunoglobulin-kappa;
            processed gene.
 SOURCE
            Mouse (BALB/c) mature B cell, cDNA to mRNA, clone AN12K.
  ORGANISM
            Mus Ausculus
            Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
            Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
            1 (bases 1 to 56)
  AUTHORS
            Rule, G.S.
  JOURNAL
            Unpublished (1988) Stanford U., Chem. Dept., Stanford CA 94305.
  STANDARD full automatic
 REFERENCE
            2 (bases 1 to 456)
  AUTHORS
            Leahy, D.J., Rule, G.S., Whittaker, M.M. and McConnell, H.M.
  TITLE
            Sequence of 12 monoclonal anti-dinitrophenyl spin-label anti-bodies
            for NMR studies
            Proc. Natl. Acad. Sci. U.S.A. 85, 3661-3665 (1988)
   JOURNAL
  STANDARD
            full automatic
 COMMENT
            Draft entry and computer-readable sequence [2] kindly submitted by
            G.Rule, 20-JUL-1988.
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Residue Identity =
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                           120
                                    130
                                              140
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200

210

220

230

240

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13. US-07-977-702-2 (1-318)
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                M.musculus NPCRLB1 mRNA for immunoglobulin light c
     MMNPCRLB1 standard; RNA; ROD; 321 BP.
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 AC
     X70095; X51612;
 XX
     19-MAR-1993 (Rel. 35, Created)
 DT
 DT
     19-MAR-1993 (Rel. 35, Last updated, Version 1)
 XX
 DE
     M.musculus NPCRLB1 mRNA for immunoglobulin light chain, variable
 DE
     region
 XX
 KW
     immunoglobulin light chain variable region.
 XX
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     Mus ausculus (aouse)
 OC.
     Eukaryota; Animalia; Metazoa; Chordata; Vertebrata; Mammalia;
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     Theria; Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 XX
 RN
     [1]
 RP
     1-321
 RA
     Izui S.;
 RT
 RL
     Submitted (05-FEB-1993) on tape to the EMBL Data Library by:
 RL
     S. Izui, Dept. of Pathology, Centre Medical Universitaire, 1, Rue
 RL
     Michel Servet, CH-1211, geneva 4, SWITZERLAND
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     *source: cell_type=hybridoma;
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     *source: cell_line=CB1;
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     *source: is_geraline=N;
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Initial Score
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                                              273 Significance = 21.50
Residue Identity =
                     85% Matches
                                              273 Mismatches
                                                                    45
Gaps
                      O Conservative Substitutions
                                                                     0
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                              30
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LOCUS
           MUSICKVN
                       2002 bp ds-DNA
                                                ROD
                                                          30-JUN-1987
DEFINITION Mouse Ig germline kappa-chain V-region gene V-Ser.
 ACCESSION
           M14361
 KEYWORDS
           constant region; germline; immunoglobulin-kappa; variable region.
SOURCE
           Mouse (C.C58) liver DNA, clones EMC58-[5,7,10].
  ORGANISM
           Mus musculus
           Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
           Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
REFERENCE
           1 (bases 1 to 2002)
  AUTHORS
           Boyd, R.T., Goldrick, M.M. and Gottlieb, P.D.
           Structural differences in a single gene encoding the V-k-Ser group
  TITLE
           of light chains explain the existence of two mouse light-chain
           genetic markers
  JOURNAL
           Proc. Natl. Acad. Sci. U.S.A. 83, 9134-9138 (1986)
  STANDARD
           full automatic
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                   LAVYFCQQHYSSP"
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Initial Score
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                                               Significance = 21.41
Residue Identity =
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15. US-07-977-702-2 (1-318)
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               Mouse Ig kappa-chain mRNA V-region (VJ), from hybr
LOCUS
           MUSIGKCKP
                        341 bp ss-mRNA
                                               ROD
                                                       15-MAR-1989
DEFINITION
           Mouse Ig kappa-chain mRNA V-region (VJ), from hybridoma
           A003-40/5G7k.
ACCESSION
           M17161 J02815
KEYWORDS
           immunoglobulin light chain; immunoglobulin-kappa; joining region;
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variable region.

/partial

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nouse (BALB/C) anti-bloodgroup a nybridoma, cuma to AKNA, clone
            A003-40/5G7k.
  ORGANISM Mus musculus
            Eukaryota; Animalia; Chordata; Vertebrata; Mammalia; Theria;
            Eutheria; Rodentia; Myomorpha; Muridae; Murinae.
 REFERENCE
            1 (bases 1 to 341)
  AUTHORS
            Chen, H.-T., Kabat, E.A., Lundblad, A. and Ratcliffe, R.M.
  TITLE
            Nucleotide and translated amino acid sequences of cDNA coding for
            the variable regions of the light and heavy chains of mouse
            hybridoma antibodies to blood group A and B substances
  JOURNAL
            J. Biol. Chem. 262, 13579-13583 (1987)
  STANDARD
            full automatic
            Draft entry and printed copy of sequence for [1] kindly provided by
 COMMENT
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                                               272 Significance = 21.32
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